	FILE 'REGISTRY' ENTERED AT 11:23:10 ON 01 JUN 2010
L1	1 S ABAPERIDONE/CN
L2	10 S BELAPERIDONE/CN OR CLOZAPINE/CN OR ILOPERIDONE/CN OR OLANZAPI
L3	1 S SERTINDOLE/CN
	FILE 'HCAPLUS' ENTERED AT 11:25:32 ON 01 JUN 2010
L4	15617 S L1/THU OR L2/THU OR L3/THU OR ANTIPSYCHOTIC
L5	99986 S SSRI OR SEROTONIN OR ANTIDEPRESSANT
L6	2906 S L4 AND L5
L7	104387 S DEPRESSION OR MDD OR DEPRESSIVE
L8	843 S L6 AND L7
L9	8483 S L1/THU OR L2/THU OR L3/THU OR (ATYPICAL ANTIPSYCHOTIC) OR ARI
L10	1518 S L5 AND L9
L11	457 S L7 AND L10
L12	89 S L11 AND (PY<2003 OR AY<2003 OR PRY<2003)
L13	9038 S SUICIDE OR SUICIDAL OR SUICIDALITY
L14	5 S L12 AND L13
	FILE 'STNGUIDE' ENTERED AT 11:27:40 ON 01 JUN 2010
	FILE 'HCAPLUS' ENTERED AT 11:28:34 ON 01 JUN 2010
L15	62 S L11 AND (PY<2002 OR AY<2002 OR PRY<2002)
L16	2105 S DOPAMINE AND D4
L17	219 S L9 AND L16
L18	101 S L17 AND (PY<2002 OR AY<2002 OR PRY<2002)
L19	1775 S DOPAMINE (3A) D4
L20	1779 S L9 AND L19
L21	75 S L20 AND (PY<2002 OR AY<2002 OR PRY<2002)
ь21 L22	115310 S DEPRESSION OR ANTIDEPRESSANT
ьzz L23	
ムムろ	3 S L21 AND L22

=> file reg
COST IN U.S. DOLLARS

FULL ESTIMATED COST

SINCE FILE TOTAL
ENTRY SESSION
0.22 0.22

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Property values tagged with IC are from the ${\tt ZIC/VINITI}$ data file provided by InfoChem.

STRUCTURE FILE UPDATES: 31 MAY 2010 HIGHEST RN 1226488-46-5 DICTIONARY FILE UPDATES: 31 MAY 2010 HIGHEST RN 1226488-46-5

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TSCA INFORMATION NOW CURRENT THROUGH January 8, 2010.

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http://www.cas.org/support/stngen/stndoc/properties.html

=> s abaperidone/cn

L1 1 ABAPERIDONE/CN

=> s belaperidone/cn or clozapine/cn or iloperidone/cn or olanzapine/cn or perospirone/cn or risperidone/cn or sertindone/cn or tiospirone/cn or ziprasidone/cn or zotepine/cn or quetiapine/cn or blonaserin/cn

1 BELAPERIDONE/CN

1 CLOZAPINE/CN

1 ILOPERIDONE/CN

1 OLANZAPINE/CN

1 PEROSPIRONE/CN

1 RISPERIDONE/CN

0 SERTINDONE/CN

1 TIOSPIRONE/CN

1 ZIPRASIDONE/CN

1 ZOTEPINE/CN

1 QUETIAPINE/CN

0 BLONASERIN/CN

10 BELAPERIDONE/CN OR CLOZAPINE/CN OR ILOPERIDONE/CN OR OLANZAPINE/
CN OR PEROSPIRONE/CN OR RISPERIDONE/CN OR SERTINDONE/CN OR TIOSP
IRONE/CN OR ZIPRASIDONE/CN OR ZOTEPINE/CN OR QUETIAPINE/CN OR
BLONASERIN/CN

=> s sertindole/cn

L3 1 SERTINDOLE/CN

=> file hcaplus

L2

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 78.96 79.18

FULL ESTIMATED COST

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FILE COVERS 1907 - 1 Jun 2010 VOL 152 ISS 23
FILE LAST UPDATED: 31 May 2010 (20100531/ED)
REVISED CLASS FIELDS (/NCL) LAST RELOADED: Apr 2010
USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Apr 2010

HCAplus now includes complete International Patent Classification (IPC) reclassification data for the second quarter of 2010.

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http://www.cas.org/legal/infopolicy.html

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 11/thu or 12/thu or 13/thu or antipsychotic 12 L1 1247350 THU/RL 10 L1/THU (L1 (L) THU/RL)9782 L2 1247350 THU/RL 6573 L2/THU (L2 (L) THU/RL)444 L3 1247350 THU/RL 344 L3/THU (L3 (L) THU/RL)13181 ANTIPSYCHOTIC 15617 L1/THU OR L2/THU OR L3/THU OR ANTIPSYCHOTIC T.4 => s SSRI or serotonin or antidepressant 2313 SSRI 80585 SEROTONIN 25756 ANTIDEPRESSANT L_5 99986 SSRI OR SEROTONIN OR ANTIDEPRESSANT => s 14 and 15 2906 L4 AND L5 => s depression or MDD or depressive

=> s depression or MDD or depressive
99682 DEPRESSION
1187 MDD
11633 DEPRESSIVE
L7 104387 DEPRESSION OR MDD OR DEPRESSIVE

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=> s 16 and 17
1.8
          843 L6 AND L7
=> s 11/thu or 12/thu or 13/thu or (atypical antipsychotic) or aripiprazole
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       1247350 THU/RL
            10 L1/THU
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          9782 L2
       1247350 THU/RL
          6573 L2/THU
                (L2 (L) THU/RL)
           444 L3
       1247350 THU/RL
           344 L3/THU
                (L3 (L) THU/RL)
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L9
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=> s 17 and 110
          457 L7 AND L10
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       4002027 PRY<2003
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          7984 SUICIDE
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L13
          9038 SUICIDE OR SUICIDAL OR SUICIDALITY
=> s 112 and 113
             5 L12 AND L13
T.14
=> d 114 1-5 ti abs bib
L14 ANSWER 1 OF 5 HCAPLUS COPYRIGHT 2010 ACS on STN
    Method of using a cyclooxygenase 2 (COX-2) inhibitor and a 5-HT1A receptor
     modulator as a combination therapy for pain, inflammation, and other
     conditions
AΒ
     Compns. and methods to treat or prevent pain, inflammation, or
     inflammation-related disorder, as well as a neurol. disorder involving
     neurodegeneration involve a combination of a COX-2 inhibitor and a 5-HT1A
     receptor modulator.
ΑN
     2004:452952 HCAPLUS <<LOGINID::20100601>>
DN
    141:1296
    Method of using a cyclooxygenase 2 (COX-2) inhibitor and a 5-HT1A receptor
     modulator as a combination therapy for pain, inflammation, and other
     conditions
```

```
PA
     Pharmacia Corporation, USA
SO
     PCT Int. Appl., 195 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 1
                                        APPLICATION NO. DATE
     PATENT NO.
                        KIND DATE
                        ____
                                            _____
     WO 2004045509 A2 20040603 WO 2003-US35739 20031111 <--
WO 2004045509 A3 20040826
PΙ
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             CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE,
             GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK,
             LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ,
             OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM,
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US 20040147581 A1 20040729 US 2003-702403 20031105 <--
AU 2003295431 A1 20040615 AU 2003-295431 20031111 <--
PRAI US 2002-427198P P 20021118 <--
WO 2003-US35739 W 20031111
OSC.G 13
RE.CNT 2
              THERE ARE 13 CAPLUS RECORDS THAT CITE THIS RECORD (16 CITINGS)
              THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L14 ANSWER 2 OF 5 HCAPLUS COPYRIGHT 2010 ACS on STN
ТΤ
     Combination therapy for depression, prevention of
     suicide, and various medical and psychiatric conditions
AΒ
     The present invention relates to a new method of treatment for persons
     meeting diagnoses for major depressive disorder, or other
     unipolar (non-bipolar, nonpsychotic and non-treatment resistant)
     depression. The method comprises administering a combination of
     two categories of drugs, antipsychotics or dopamine system stabilizers, in
     combination with a newer antidepressant such as a selective
     serotonin reuptake inhibitor, as initial treatment or as soon as
     possible. The method targets the prevention of suicide, and
     provides other benefits including preventing disease progression
     development of tolerance toward the antidepressants. Another aspect of
     the invention relates to using the method for alleviating cognitive
     distortion and related functional impairment or health risks, and/or using
     the method for smoking cessation or nicotine withdrawal.
     2004:100942 HCAPLUS <<LOGINID::20100601>>
ΑN
DN
     140:139528
     Combination therapy for depression, prevention of
ΤI
     suicide, and various medical and psychiatric conditions
IN
     Migaly, Peter
PA
     USA
SO
     PCT Int. Appl., 28 pp.
     CODEN: PIXXD2
DT
     Patent
     English
LA
FAN.CNT 1
     PATENT NO.
                        KIND DATE APPLICATION NO. DATE
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      WO 2004010932
      A2
      20040205

      WO 2004010932
      A3
      20040722

PΙ
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Stephenson, Diane T.; Taylor, Duncan P.

TN

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CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
             PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ,
             UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
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             FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
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     CA 2529857
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                         Α1
     AU 2003268026
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     US 20040204401
                                20041014
                                            US 2003-627358
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                          Α1
     EP 1551393
                          Α2
                                20050713
                                            EP 2003-748977
                                                                   20030725 <--
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             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
                                20050819
     MX 2005000294
                                         MX 2005-294
                         Α
                                                                   20050104 <--
PRAI US 2002-319436P
                          Ρ
                                20020730 <--
     US 2003-627358
                                20030725
                          Α
     WO 2003-US23326
                          W
                                20030725
OSC.G
              THERE ARE 5 CAPLUS RECORDS THAT CITE THIS RECORD (5 CITINGS)
RE.CNT 2
              THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L14 ANSWER 3 OF 5 HCAPLUS COPYRIGHT 2010 ACS on STN
     Putting metabolic side effects into perspective: Risks versus benefits of
     atypical antipsychotics
     A review. The lengthy list of the side effects and morbidity associated with
AB
     the atypical antipsychotics might make a patient with psychosis and his or
     her caregivers so concerned about the use of any of these medications,
     particularly those associated with a higher risk of diabetes, weight gain, or
     increased lipid levels, that they would prefer to avoid all of them.
     However, schizophrenia is associated with a relatively high risk for several
     diseases, including diabetes, that is independent of the risks that are
     linked to atypical antipsychotic use. Therefore, the
     clinician who might think, "Why use atypicals if using the typical drugs
     will escape the problems of monitoring and all the associated effects of
     diabetes and hyperglycemia" needs to know that these problems cannot be
     avoided simply by choosing typical antipsychotics. Clinicians, patients,
     and concerned family members must balance the significant benefits of
     atypical antipsychotic treatment - improved cognition,
     reduced suicidality, and less depression - against the
     risks of metabolic disturbances and select a course of treatment that
     includes a realistic monitoring program.
ΑN
     2002:75124 HCAPLUS <<LOGINID::20100601>>
DΝ
     136:272542
     Putting metabolic side effects into perspective: Risks versus benefits of
ΤI
     atypical antipsychotics
     Meltzer, Herbert Y.
ΑU
     Department of Psychiatry and Pharmacology, Division of Psychopharmacology,
CS
     Vanderbilt University Medical Center, Nashville, TN, 37212, USA
SO
     Journal of Clinical Psychiatry (2001), 62(Suppl. 27), 35-39
     CODEN: JCLPDE; ISSN: 0160-6689
PΒ
     Physicians Postgraduate Press, Inc.
     Journal; General Review
DT
     English
LA
              THERE ARE 6 CAPLUS RECORDS THAT CITE THIS RECORD (6 CITINGS)
OSC.G
        6
RE.CNT
       47
              THERE ARE 47 CITED REFERENCES AVAILABLE FOR THIS RECORD
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L14 ANSWER 4 OF 5 HCAPLUS COPYRIGHT 2010 ACS on STN

ALL CITATIONS AVAILABLE IN THE RE FORMAT

TI Treatment of suicidality in schizophrenia

- A review with 48 refs. Between 4 and 13% of people with schizophrenia AΒ commit suicide and between 25 and 50% make a suicide attempt, a reflection of the devastating toll this syndrome takes on the quality of life, i.e., the subjective and objective sense of well-being. Many risk factors for suicide in schizophrenia have been identified, the most important of which are previous suicide attempts, depression, hopelessness, substance abuse, and male gender. Insight into having a serious mental illness and less severe cognitive impairment are also associated with increased risk for suicide in schizophrenia, most likely when accompanied by feelings of hopelessness. Typical neuroleptic drugs have not been shown to reduce the risk of suicide. However, several types of evidence suggest that clozapine, an atypical antipsychotic drug, appreciably reduces the suicide attempt and completion rates in schizophrenia and schizo-affective disorder, perhaps by as much as 75-85%. Other atypical antipsychotic drugs may have a similar effect, but direct evidence is lacking. Improvement in pos. and neg. symptoms, reduced extrapyramidal side effects (EPS), a direct antidepressant action, improved cognitive function, and improved compliance may contribute to reduced suicidality. The International Suicide Prevention Trial (InterSePT) is a large prospective, randomized study intended to compare the effectiveness of clozapine with that of olanzapine in reducing suicide and suicide-related events in schizophrenic and schizoaffective patients. Some information about suicidality in the patient sample is reported here.
- AN 2001:480353 HCAPLUS <<LOGINID::20100601>>
- DN 135:266558
- TI Treatment of suicidality in schizophrenia
- AU Meltzer, Herbert Y.
- CS Division of Psychopharmacology, Vanderbilt University School of Medicine, Nashville, TN, 37212, USA
- SO Annals of the New York Academy of Sciences (2001), 932(Clinical Science of Suicide Prevention), 44-60 CODEN: ANYAA9; ISSN: 0077-8923
- PB New York Academy of Sciences
- DT Journal; General Review
- LA English
- OSC.G 12 THERE ARE 12 CAPLUS RECORDS THAT CITE THIS RECORD (12 CITINGS)
- RE.CNT 48 THERE ARE 48 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L14 ANSWER 5 OF 5 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Olanzapine: Preclinical and clinical profiles of a novel antipsychotic agent
- AΒ A review with 278 refs. The novel antipsychotic agent olanzapine (Zyprexa, Eli Lilly and Company) is a thienobenzodiazepine analog marketed for the treatment of schizophrenia. Olanzapine's diverse receptor binding profile and greater affinity for serotonin receptors over dopamine receptors is thought to impart antipsychotic efficacy with a low incidence of serious extrapyramidal symptoms (EPS). With once daily dosing steady-state plasma concns. reached within approx. 1 wk. Olanzapine is extensively metabolized by the liver, is mostly excreted in the urine, and has few drug intractions. In clin. trials, the efficacy of olanzapine for treating schizophrenia is better than placebo and haloperidol and comparable to risperidone. Olanzapine may also ameliorate some comorbid symptoms including neg. symptoms, depression, anxiety, substance abuse, and cognitive dysfunction, and it is effective in the long-term maintenance of response, treatment-resistance, and improving quality of life. The overall direct costs are lower with olanzapine treatment compared with haloperidol or risperidone treatment.

In clin. trials, olanzapine demonstrates a favorable safety profile. The most frequently reported treatment-emergent adverse events are somnolence, schizophrenic reaction, insomnia, headache, agitation, rhinitis, and weight gain. Significantly fewer EPS (based on formal rating scales) and incidences of tardive dyskinesia have been reported for olanzapine compared with haloperidol. Olanzapine has not been associated with persistent elevations of prolactin above the upper limit of normal nor has it been associated with clin. significant changes in cardiac QTc interval. Fewer incidences of suicide attempts have been reported with olanzapine compared with placebo, haloperidol, or risperidone treatments. There is evidence that olanzapine may be effective in the treatment of mood disorders, psychosis associated with. Alzheimer's disease, obsessive-compulsive disorder, pervasive developmental disorders, and delirium. Patients with schizophrenia have been successfully switched from other antipsychotics to olanzapine. In conclusion, olanzapine offers a significantly improved risk-to-benefit profile compared with haloperidol and possibly risperidone, and thus should be considered an important treatment option for schizophrenia and related disorders.

AN 2001:49031 HCAPLUS <<LOGINID::20100601>>

DN 135:86379

- TI Olanzapine: Preclinical and clinical profiles of a novel antipsychotic agent
- AU Tollefson, Gary D.; Taylor, Cindy C.
- CS Lilly Research Laboratories Lilly Corporate Center, Eli Lilly and Company, Indianapolis, IN, 46285, USA
- SO CNS Drug Reviews (2000), 6(4), 303-363 CODEN: CDREFB; ISSN: 1080-563X
- PB Neva Press
- DT Journal; General Review
- LA English
- OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)
- RE.CNT 278 THERE ARE 278 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> file stnguide COST IN U.S. DOLLARS	SINCE FILE	TOTAL
FULL ESTIMATED COST	ENTRY 27.14	SESSION 106.32
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE ENTRY	TOTAL SESSION
CA SUBSCRIBER PRICE	-4.25	-4.25

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FILE CONTAINS CURRENT INFORMATION.
LAST RELOADED: May 28, 2010 (20100528/UP).

=> file hcaplus COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 0.14 106.46 DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL SESSION ENTRY CA SUBSCRIBER PRICE 0.00 -4.25

FILE 'HCAPLUS' ENTERED AT 11:28:34 ON 01 JUN 2010

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FILE COVERS 1907 - 1 Jun 2010 VOL 152 ISS 23
FILE LAST UPDATED: 31 May 2010 (20100531/ED)
REVISED CLASS FIELDS (/NCL) LAST RELOADED: Apr 2010
USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Apr 2010

HCAplus now includes complete International Patent Classification (IPC) reclassification data for the second quarter of 2010.

CAS Information Use Policies apply and are available at:

http://www.cas.org/legal/infopolicy.html

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s l11 and (PY<2002 or AY<2002 or PRY<2002) 22006893 PY<2002 4244485 AY<2002 3713094 PRY<2002

L15 62 L11 AND (PY<2002 OR AY<2002 OR PRY<2002)

=> d 115 1-62 ti abs bib

- L15 ANSWER 1 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Methods and compositions using cyclooxygenase 2 (COX-2) inhibitors for the treatment of psychiatric disorders, and combination therapies
- AB A method for the prevention, treatment, or inhibition of a psychiatric disorder, in particular schizophrenia, is described which comprises administering a COX-2 inhibitor or prodrug thereof to a subject. Moreover, a method for the prevention, treatment, or inhibition of a psychiatric disorder, in particular schizophrenia or depressive disorders, is disclosed comprising administering to a subject a COX-2 inhibitor or prodrug thereof in combination with a neuroleptic drug or an antidepressant. Compns. and kits that are suitable for the practice of the method are also described.
- AN 2006:740188 HCAPLUS <<LOGINID::20100601>>
- DN 145:159849
- TI Methods and compositions using cyclooxygenase 2 (COX-2) inhibitors for the treatment of psychiatric disorders, and combination therapies
- IN Muller, Norbert
- PA Germany
- SO U.S. Pat. Appl. Publ., 34 pp., Cont.-in-part of U.S. Ser. No. 157,969. CODEN: USXXCO
- DT Patent
- LA English

FAN.CNT 3

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PATENT NO.
                         KIND DATE APPLICATION NO. DATE
      US 20060167074 A1 20060727 US 2005-320757 20051230 <--
US 20030130334 A1 20030710 US 2002-157969 20020531 <--
EP 1627639 A2 20060222 EP 2005-24864 20020531 <--
PΤ

      US
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      1627639
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      EP
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      20091223

            R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
                   IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
       JP 2008297308 A 20081211 JP 2008-188890 20080722 <--
PRAI DE 2001-10129328
      DE 2001-10129328 A 20010619 <--
US 2002-364904P P 20020314
US 2002-157969 A2 20020531
DE 2001-10129320 A 20010619 <--
EP 2002-738138 A3 20020531
JP 2003-504886 A3 20020531
                                   A
                                            20010619 <--
      MARPAT 145:159849
OS
L15 ANSWER 2 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
       Combination therapy for treatment of refractory depression
TΤ
AΒ
       The invention provides methods and compns. for the treatment of
       depressive states refractory to treatment with traditional anti-
depressive therapies alone. These methods and compns. employ a
       compound having activity as an atypical antipsychotic
       and a serotonin reuptake inhibitor. This invention also
       provides methods of providing rapid onset treatments of major
       depression which employing a compound having activity as an
       atypical antipsychotic and a serotonin
       reuptake inhibitor.
AN
      2003:892442 HCAPLUS <<LOGINID::20100601>>
DN
      139:345944
ΤI
      Combination therapy for treatment of refractory depression
      Tollefson, Gary Dennis
ΙN
PA
       Eli Lilly and Company, USA
SO
      U.S. Pat. Appl. Publ., 10 pp.
       CODEN: USXXCO
DT
      Patent
      English
LA
FAN.CNT 2
      PATENT NO. KIND DATE APPLICATION NO. DATE
      US 20030212060 A1 20031113 US 2002-144159 20020510 <--
US 6960577 B2 20051101
CA 2332814 A1 19991202 CA 1999-2332814 19990521 <--
CA 2332814 C 20081104
WO 9961027 A1 19991202 WO 1999-US11276 19990521 <--
PΙ
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HR 2000000797 A2 20011031 HR 2000-797 20001120 <--
NO 2000005885 A 20010117 NO 2000-5885 20001121 <--
HK 1040055 A1 20050401 HK 2002-101563 20020228 <--
PRAI US 1998-86444P P 19980522 <--
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US 2000-700254 B1 20001109 <--
OSC.G 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD (3 CITINGS)
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- L15 ANSWER 3 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Methods and compositions using a cyclooxygenase 2 (COX-2) inhibitor for the treatment of psychiatric disorders
- AB A method for the prevention, treatment, or inhibition of a psychiatric disorder, in particular schizophrenia, is described which comprises administering a COX-2 inhibitor, or prodrug thereof, to a subject. Moreover, a method for the prevention, treatment, or inhibition of a psychiatric disorder, in particular schizophrenia or a depressive disorder, is disclosed, comprising administering to a subject a COX-2 inhibitor or prodrug thereof in combination with a neuroleptic drug or an antidepressant. Compns. and kits that are suitable for the practice of the method are also described.
- AN 2003:532347 HCAPLUS <<LOGINID::20100601>>
- DN 139:79173
- TI Methods and compositions using a cyclooxygenase 2 (COX-2) inhibitor for the treatment of psychiatric disorders
- IN Muller, Norbert
- PA Germany
- SO U.S. Pat. Appl. Publ., 27 pp. CODEN: USXXCO
- DT Patent
- LA English
- FAN.CNT 3

	PATENT NO.						IND DATE			i	APPI	ICAT	DATE							
ΡI	US	US 20030130334			A1	1 20030710			1	 JS 2	2002-	2	<							
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	JΡ	2003	-504	886		А3		2002	0531											
	US	2002	-157	969		A2		2002	0531											
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- L15 ANSWER 4 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Methods for treating depression and other CNS disorders using enantiomerically enriched desmethyl- and didesmethyl- metabolites of citalogram

AΒ This invention relates to the preparation of I and II and derivs. of I and II in their racemic, enantiomerically enriched, or optically pure forms. This invention further relates to novel compns. of matter containing enantiomerically enriched (-)-desmethylcitalogram (-)-III (R = Me), (+)-didesmethylcitalopram (+)-III (R = Me), or (-)-didesmethylcitalopram (-)-III (R = H) or mixts. thereof in optimal ratios. Contrary to prior teachings, the enantiomerically enriched citalopram metabolites disclosed herein possess potent serotonin reuptake inhibitory activity, with minimal inhibitory effects on the reuptake of other known monoamines, e.g., norepinephrine (NE) or dopamine (DA). For example, stepwise reaction of 1-oxo-1,3-dihydroisobenzofuran-5-carbonitrile with 4-fluorophenylmagnesium bromide and the chiral Grignard reagent, which was prepared from 2-(2-bromoethy1)-[1,3]dioxolane and Mg powder, in THF gave II.Cyclization using mesyl chloride in CH2Cl2, followed by reduction provided the I. Reaction of the aldehyde with (-)-tert-butylsulfinamide in the presence of Ti(OEt)4 in EtOH afforded the sulfinamide, which was reduced to the amine III (R = H) with 10% HCl in MeOH. Protection of the amine with BOC anhydride in the presence of TEA in CH2Cl2 provided the enantiomerically enriched isomers, which were separated on a chiral column and subsequently deprotected with TFA to give (+)-III (R = H) and (-)-III (R = H)H). In biol. assays, (-)-III (R = H) and (+)-III (R = H) strongly inhibited serotonergic 5-HT receptor activity with Ki values of 5.8 nM and 90 nM, resp., with little effect on NE and DA transporter activity. By comparison, racemic citalogram inhibited serotonin reuptake with a Ki of 3.9 nM. The present invention also discloses methods for treating disorders, dysfunctions and diseases for which inhibition of serotonin reuptake is therapeutically beneficial. In particular, the present invention discloses a method for treating various forms of depression and other CNS disorders with pharmaceutical compns. described herein.

AN 2003:376842 HCAPLUS <<LOGINID::20100601>>

DN 138:385297

TI Methods for treating depression and other CNS disorders using

enantiomerically enriched desmethyl- and didesmethyl- metabolites of citalopram

- Bush, Larry R.; Currie, Mark G.; Senanayake, Chris H.; Fang, Kevin Q. ΙN
- PΑ Sepracor, Inc., USA
- SO PCT Int. Appl., 58 pp.

CODEN: PIXXD2

DTPatent

LA English

FAN.	CNT	1						
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PI		2003040121 W: AE, AG, AL, CO, CR, CU, GM, HR, HU, LS, LT, LU, PL, PT, RO,	A1 20030515 AM, AT, AU, AZ, CZ, DE, DK, DM, ID, IL, IN, IS, LV, MA, MD, MG,	WO 2002-US35408 BA, BB, BG, BR, BY, BZ, DZ, EC, EE, ES, FI, GB, JP, KE, KG, KP, KR, KZ, MK, MN, MW, MX, MZ, NO, SI, SK, SL, TJ, TM, TN,	20021105 < CA, CH, CN, GD, GE, GH, LC, LK, LR, NZ, OM, PH,			
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	3	1 THERE ARE	1 CAPLUS RECORDS	THAT CITE THIS RECORD ES AVAILABLE FOR THIS RE	•			

- L15 ANSWER 5 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- Use of cyclooxygenase 2 (COX-2) inhibitors for the treatment of ΤI schizophrenia, delusional disorders, affective disorders, autism, or tic disorders

ALL CITATIONS AVAILABLE IN THE RE FORMAT

- AΒ The invention discloses the use of a COX-2 inhibitor for the treatment of psychiatric disorders, e.g. schizophrenia, delusional disorders, affective disorders, autism or tic disorders, in particular chronic schizophrenic psychoses and schizoaffective psychoses, temporary acute psychotic disorders, depressive episodes, recurring depressive episodes, manic episodes and bipolar affective disorders. Moreover, the invention discloses the use of a COX-2 inhibitor, in particular celecoxib, in combination with a neuroleptic drug, in particular risperidone, or an antidepressant, for the treatment of psychiatric disorders such as schizophrenia, delusional disorders, affective disorders, autism or tic disorders.
- ΔN 2002:977588 HCAPLUS <<LOGINID::20100601>>

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138:33362
DN
     Use of cyclooxygenase 2 (COX-2) inhibitors for the treatment of
ΤI
     schizophrenia, delusional disorders, affective disorders, autism, or tic
     disorders
     Muller, Norbert
IN
PA
     Germany
SO
     PCT Int. Appl., 58 pp.
     CODEN: PIXXD2
DT
     Patent
     English
LA
FAN.CNT 3
                         KIND DATE
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     WO 2002102297 A2 20021227 WO 2002-EP6013
WO 2002102297 A3 20030501
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A3 20020531
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               THERE ARE 5 CAPLUS RECORDS THAT CITE THIS RECORD (5 CITINGS)
OSC.G 5
RE.CNT 2
               THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
               ALL CITATIONS AVAILABLE IN THE RE FORMAT
L15 ANSWER 6 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
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Atypical antipsychotic-antidepressant

combination for treatment of depression, obsessive compulsive disorder, and psychosis

AB The invention provides a method for treating depression, obsessive compulsive disorder, and psychosis in a mammal, including a human, by administering to the mammal an atypical antipsychotic in combination with an antidepressant agent with improvement in efficiency. It also provides pharmaceutical compns. containing a pharmaceutically acceptable carrier, an atypical antipsychotic, and a serotonin reuptake inhibitor.

AN 2002:674788 HCAPLUS <<LOGINID::20100601>>

DN 137:195595

TI Atypical antipsychotic-antidepressant combination for treatment of depression, obsessive compulsive disorder, and psychosis

IN Howard, Harry R., Jr.

PA Pfizer Inc., USA

SO U.S. Pat. Appl. Publ., 20 pp. CODEN: USXXCO

DT Patent

LA English

FAN.CNT 1

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- L15 ANSWER 7 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
 TI Pharmaceuticals containing a combination of norepinephrine reuptake inhibitors and neuroleptics
- AB A composition comprising: (a) a pharmaceutically effective amount of one or more

norepinephrine reuptake inhibitors or a salt; and (b) 1 or more neuroleptics is provided. The composition is useful in treating disorders or diseases of the central nervous system, and particularly useful in treating schizophrenia. A pharmaceutical composition was prepared by combining reboxetine with a neuroleptic in an acceptable carrier. The composition contains 0.01-10 mg rebexetine and 25-300 mg clozapine.

AN 2002:521465 HCAPLUS <<LOGINID::20100601>>

DN 137:98994

- TI Pharmaceuticals containing a combination of norepinephrine reuptake inhibitors and neuroleptics
- IN Wong, Erik Ho Fong; Gallen, Christopher C.; Svensson, Torgny

PA Pharmacia & Upjohn Company, USA; Pharmacia AB

SO PCT Int. Appl., 22 pp. CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PATENT NO. KIND DATE APPLICATION NO. DATE

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     US 2001-35100
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              THERE ARE 12 CAPLUS RECORDS THAT CITE THIS RECORD (12 CITINGS)
RE.CNT 6
              THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L15 ANSWER 8 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
     (+)-1-(3,4-dichlorophenyl)-3-azabicyclo[3.1.0]hexane, compositions
ΤI
     thereof, and uses as an anti-depressant agent
AΒ
     The present invention relates to (+)-1-(3,4-dichlorophenyl)-3-
     azabicyclo[3.1.0]hexane and pharmaceutically acceptable salts thereof,
     compns. comprising (+)-1-(3,4-dichlorophenyl)-3-azabicyclo[3.1.0]hexane or
     a pharmaceutically acceptable salt thereof, and methods for treating or
     preventing depression in a patient comprising administering
     (+)-1-(3,4-dichlorophenyl)-3-azabicyclo[3.1.0]hexane or a pharmaceutically
     acceptable salt thereof. The (+)-1-(3,4-dichloropheny1)-3-
     azabicyclo[3.1.0]hexane or pharmaceutically acceptable salt thereof is
     preferably substantially free of its corresponding (-)-enantiomer. The +
     isomer is obtained by HPLC resolution on a CHIRALPAK AD column. The + isomer
     has greater affinity for both norepinephrine and serotonin
     uptake sites in rat forebrain membranes than the ± compound The + isomer
     is administered along with a known antidepressant, anxiolytic,
     antipsychotic or antiobesity agent in treatment of various
     depression conditions including depression associated with
     anxiety, seizures, menopause, alcoholism, etc.
     2002:290820 HCAPLUS <<LOGINID::20100601>>
ΑN
DN
     136:304102
     (+)-1-(3,4-dichlorophenyl)-3-azabicyclo[3.1.0]hexane, compositions
ΤI
     thereof, and uses as an anti-depressant agent
     Lippa, Arnold Stan; Epstein, Joseph William
ΙN
PA
     Dov Pharmaceutical, Inc., USA
SO
     U.S., 7 pp.
     CODEN: USXXAM
DT
     Patent
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LA

English

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FAN.CNT 4
                                                 KIND DATE APPLICATION NO. DATE

B1 20020416 US 2001-758883 20010111 <--

A1 20020829 CA 2002-2434616 20020111 <--
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              US 6372919
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                                    GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
                                    LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
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              AU 2002251758 A1 20020904 AU 2002-251758 20020111 <--

AU 2002251758 B2 20080103

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 RE.CNT 28
                                      THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS RECORD
                                      ALL CITATIONS AVAILABLE IN THE RE FORMAT
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- L15 ANSWER 9 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Putting metabolic side effects into perspective: Risks versus benefits of atypical antipsychotics
- AB A review. The lengthy list of the side effects and morbidity associated with the atypical antipsychotics might make a patient with psychosis and his or her caregivers so concerned about the use of any of these medications, particularly those associated with a higher risk of diabetes, weight gain, or increased lipid levels, that they would prefer to avoid all of them. However, schizophrenia is associated with a relatively high risk for several diseases, including diabetes, that is independent of the risks that are linked to atypical antipsychotic use. Therefore, the clinician who might think, "Why use atypicals if using the typical drugs will escape the problems of monitoring and all the associated effects of diabetes and hyperglycemia" needs to know that these problems cannot be

avoided simply by choosing typical antipsychotics. Clinicians, patients, and concerned family members must balance the significant benefits of atypical antipsychotic treatment - improved cognition, reduced suicidality, and less depression - against the risks of metabolic disturbances and select a course of treatment that includes a realistic monitoring program.

- AN 2002:75124 HCAPLUS <<LOGINID::20100601>>
- DN 136:272542
- TI Putting metabolic side effects into perspective: Risks versus benefits of atypical antipsychotics
- AU Meltzer, Herbert Y.
- CS Department of Psychiatry and Pharmacology, Division of Psychopharmacology, Vanderbilt University Medical Center, Nashville, TN, 37212, USA
- SO Journal of Clinical Psychiatry (2001), 62(Suppl. 27), 35-39 CODEN: JCLPDE; ISSN: 0160-6689
- PB Physicians Postgraduate Press, Inc.
- DT Journal; General Review
- LA English
- OSC.G 6 THERE ARE 6 CAPLUS RECORDS THAT CITE THIS RECORD (6 CITINGS)
- RE.CNT 47 THERE ARE 47 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 10 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Fluvoxamine as an adjunctive agent in schizophrenia
- AΒ A review. Schizophrenia is a common mental disorder that has an early onset and rates high as a cause of medical disability. Antipsychotic agents are the mainstay of treatment but response is often inadequate. Neg. symptoms (disturbances in volition, social interaction and affective functions) are particularly difficult to treat and form a major obstacle to rehabilitation. A promising approach to improve response of neg. symptoms has been to add a selective serotonin reuptake inhibitor (SSRI) antidepressant to antipsychotic treatment. This review examines evidence pertaining to the efficacy, tolerability, and safety of the SSRI fluvoxamine, combined with antipsychotic agents, in the treatment of neg. symptoms in schizophrenia. Important methodol. issues, such as differentiating primary and secondary neg. symptoms, are discussed. The balance of available evidence indicates that fluvoxamine can improve primary neq. symptoms in chronic schizophrenia patients treated with typical antipsychotics and suggests that it may also do so in some patients treated with clozapine. combination is generally safe and well tolerated although, as antipsychotic drug concns. may be elevated, attention to dose and drug monitoring should be considered appropriately. Combination with clozapine may require particular caution because of potential toxicity if serum clozapine levels rise steeply. The fluvoxamine doses effective in augmentation are lower than those usually used to treat depression Evidence regarding the use of fluvoxamine augmentation to treat phenomena, such as obsessions and aggression, which may be associated with schizophrenia, is also examined
- AN 2002:50209 HCAPLUS <<LOGINID::20100601>>
- DN 136:288416
- TI Fluvoxamine as an adjunctive agent in schizophrenia
- AU Silver, Henry
- CS Sha'ar Menashe Mental Health Center, Rappaport Faculty of Medicine, Haifa, Israel
- SO CNS Drug Reviews (2001), 7(3), 283-304 CODEN: CDREFB; ISSN: 1080-563X
- PB Neva Press
- DT Journal; General Review
- LA English
- OSC.G 21 THERE ARE 21 CAPLUS RECORDS THAT CITE THIS RECORD (21 CITINGS)

- L15 ANSWER 11 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI No effect of reboxetine on plasma concentrations of clozapine, risperidone, and their active metabolites
- AΒ The effect of reboxetine on steady-state plasma concns. of the atypical antipsychotics clozapine and risperidone was studied in 14 patients with schizophrenia or schizo-affective disorder with associated depressive symptoms. Seven patients stabilized on clozapine therapy (250-500 mg/day) and seven receiving risperidone (4-6 mg/day) were given addnl. reboxetine (8 mg/day). After 4 wk of reboxetine therapy, mean plasma concns. of clozapine, norclozapine, and risperidone active moiety (sum of concns. of risperidone and 9-hydroxyrisperidone) increased slightly but not significantly by 5%, 2%, and 10%, resp. The mean plasma clozapine/norclozapine and risperidone/9-hydroxyrisperidone ratios were not modified during reboxetine treatment. Reboxetine coadministration with either clozapine or risperidone was well tolerated. These findings indicate that reboxetine has minimal effects on the metabolism of clozapine and risperidone and may be added safely to patients receiving maintenance treatment with these two antipsychotics.
- AN 2002:823 HCAPLUS <<LOGINID::20100601>>
- DN 136:193677
- TI No effect of reboxetine on plasma concentrations of clozapine, risperidone, and their active metabolites
- AU Spina, Edoardo; Avenoso, Angela; Scordo, Maria Gabriella; Ancione, Maria; Madia, Aldo; Levita, Antonino
- CS Department of Clinical and Experimental Medicine and Pharmacology, Section of Pharmacology, University of Messina, Messina, 98125, Italy
- SO Therapeutic Drug Monitoring (2001), 23(6), 675-678 CODEN: TDMODV; ISSN: 0163-4356
- PB Lippincott Williams & Wilkins
- DT Journal
- LA English
- OSC.G 7 THERE ARE 7 CAPLUS RECORDS THAT CITE THIS RECORD (7 CITINGS)
- RE.CNT 23 THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 12 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Olanzapine: A review of its use in the treatment of bipolar I disorder
- A review. Olanzapine, a thienobenzodiazepine derivative, is a psychotropic AΒ agent that has shown efficacy in the treatment of patients with bipolar I disorder. Olanzapine has a multireceptorial binding profile including a greater affinity for serotonin 5-HT2A than for dopamine D2 receptors. Olanzapine 5 to 20 mg/day demonstrated significantly greater antimanic efficacy than placebo in two double-blind, randomized 3- or 4-wk trials of patients with bipolar I disorder of either manic or mixed episodes, with or without psychotic features. Addnl., in one of these trials, improvements in cognitive function and hostility were superior with olanzapine. In cohorts of severely depressed and rapid cycling patients, improvements in manic and depressive symptoms and in manic symptoms only, were superior with olanzapine compared with placebo. Significant improvements from baseline in symptoms of mania, depression, cognitive functioning and hostility were seen with olanzapine in a 49-wk extension phase study. In double-blind trials, olanzapine 10 mg/day appeared to have similar antimanic efficacy to oral lithium 440mg twice daily in the treatment of patients with pure mania (4-wk small study). In patients with acute manic or mixed episodes olanzapine 5 to 20 mg/day appeared to be more effective than oral valproate semisodium (divalproex sodium) 500 to 2500 mg/day (3-wk study) and at least as effective as oral haloperidol 3 to 15 mg/day (12-wk

study). Preliminary results from a large 6-wk placebo-controlled study suggest that olanzapine 5 to 20 mg/day in combination with mood stabilizers (lithium or valproate semisodium) provides effective augmentation of antimanic treatment of patients with bipolar I disorder, with benefits seen in the first week. Adverse events reported significantly more often with olanzapine than with placebo were somnolence, dry mouth, dizziness and bodyweight gain, and in comparison with valproate semisodium were somnolence, dry mouth, increased appetite and bodyweight gain. Olanzapine was generally well tolerated with no clin. relevant abnormalities in laboratory tests, vital signs or ECG results. Conclusion: Olanzapine demonstrated superior efficacy compared with placebo in the short-term treatment of patients with bipolar I disorder with manic or mixed episodes, with or without psychotic features, and was generally well tolerated. According to preliminary data the antimanic efficacy of olanzapine appears similar to that of haloperidol and better than that of valproate semisodium in patients with bipolar I disorder experiencing a manic or mixed episode; among nonpsychotic patients with manic or mixed episodes olanzapine appears to be superior to haloperidol. Available data support the choice of olanzapine as an option in the short-term management of mania in patients with bipolar I disorder with manic or mixed episodes, with or without psychotic features.

- AN 2001:934947 HCAPLUS <<LOGINID::20100601>>
- DN 136:226160
- TI Olanzapine: A review of its use in the treatment of bipolar I disorder
- AU Bhana, Nila; Perry, Caroline M.
- CS Adis International Limited, Auckland, N. Z.
- SO CNS Drugs (2001), 15(11), 871-904 CODEN: CNDREF; ISSN: 1172-7047
- PB Adis International Ltd.
- DT Journal; General Review
- LA English

ΑN

- OSC.G 14 THERE ARE 14 CAPLUS RECORDS THAT CITE THIS RECORD (14 CITINGS)
- RE.CNT 153 THERE ARE 153 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 13 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Extended Radioligand Binding Profile of Iloperidone: A Broad Spectrum Dopamine/Serotonin/Norepinephrine Receptor Antagonist for the Management of Psychotic Disorders
- AB Iloperidone is a novel psychotropic compound currently undergoing Phase III trials. Its affinity for human dopamine and 5-HT2A and 5-HT2C receptors has been reported previously. This report presents the affinity of iloperidone for a largely extended number of human neurotransmitter receptors. In a few instances human receptors were not available and receptor studies were performed on tissues from laboratory animals. The present

data indicate that iloperidone displays high affinity (KI < 10 nM) for norepinephrine $\alpha 1-$ adrenoceptors, dopamine D3 and serotonin 5-HT2A receptors. Intermediate affinity (10-100 nM) was found for norepinephrine $\alpha 2C-$ adrenoceptors, dopamine D2A and D4 receptors and serotonin 5-HT1A, 5-HT1B, 5-HT2C and 5-HT6 receptors. The affinity for all other receptors was below 100 nM, including norepinephrine $\alpha 2A$, $\alpha 2B$, $\beta 1$, and $\beta 2$, muscarine M1-M5, histamine H1, dopamine D1 and D5, CCKA and CCKB, 5-HT7, dopamine and norepinephrine transporters. Thus, iloperidone targets a selective set of dopamine, norepinephrine and serotonin receptor subtypes. The affinity for this particular set of receptors indicates that iloperidone has the potential to be a broad spectrum antipsychotic, with efficacy against pos., neg., depressive and cognitive symptoms of schizophrenia, and a low propensity to induce side effects. 2001:922467 HCAPLUS <<LOGINID::20100601>>

- DN 137:150052
- TI Extended Radioligand Binding Profile of Iloperidone: A Broad Spectrum Dopamine/Serotonin/Norepinephrine Receptor Antagonist for the Management of Psychotic Disorders
- AU Kalkman, Hans Otto; Subramanian, Natarajan; Hoyer, Daniel
- CS Novartis Pharma, Basel, Switz.
- SO Neuropsychopharmacology (2001), 25(6), 904-914 CODEN: NEROEW; ISSN: 0893-133X
- PB Elsevier Science Inc.
- DT Journal
- LA English
- OSC.G 29 THERE ARE 29 CAPLUS RECORDS THAT CITE THIS RECORD (29 CITINGS)
- RE.CNT 69 THERE ARE 69 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 14 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Risperidone safety and efficacy in the treatment of bipolar and schizoaffective disorders: Results from a 6-month, multicenter, open study
- AΒ The goal of this study was to assess the efficacy and safety of risperidone in bipolar and schizoaffective disorders. 541 Patients entered this open, multicenter, 6-mo study. Patients were entered provided that they fulfilled DSM-IV criteria for bipolar disorder or schizoaffective disorder, bipolar type, during a manic, hypomanic, mixed, or depressive episode. Risperidone was added to any previous mood-stabilizing medication that the patients were taking. Efficacy was assessed with the Young Mania Rating Scale (YMRS), the Hamilton Rating Scale for Depression (HAM-D), the Pos. and Neg. Syndrome Scale $\,$ (PANSS), and the Clin. Global Impressions scale (CGI). Extrapyramidal symptoms (EPS) were assessed using the UKU Side Effect Rating Scale. 430 Patients completed the study. Addition of risperidone produced highly significant improvements (p < .0001) on the YMRS and HAM-D at both 6 wk and 6 mo and on the CGI and the scales of the PANSS at both $4\ \mathrm{wk}$ and $6\ \mathrm{mo}$. There was a significant reduction in UKU total and subscale scores at 6 mo. The mean dose of risperidone was 3.9 mg/day. There was no single case of new-emergent tardive dyskinesia, and there was a very low incidence of exacerbation of mania within the first 6 wk (2%). Adverse events were few and mostly mild, the most frequent being EPS and weight gain. This large study provides addnl. evidence that risperidone is effective and well tolerated when combined with mood stabilizers in the treatment of bipolar disorder and schizoaffective disorder, bipolar type. Previous concerns about exacerbation of manic symptoms were not confirmed.
- AN 2001:912080 HCAPLUS <<LOGINID::20100601>>
- DN 136:177875
- TI Risperidone safety and efficacy in the treatment of bipolar and schizoaffective disorders: Results from a 6-month, multicenter, open study
- AU Vieta, Eduard; Goikolea, Jose M.; Corbella, Barbara; Benabarre, Antonio; Reinares, Maria; Martinez, Guadalupe; Fernandez, Antonio; Colom, Francesc; Martinez-Aran, Anabel; Torrent, Carla
- CS Department of Psychiatry, University of Barcelona, Barcelona, 08036, Spain
- SO Journal of Clinical Psychiatry (2001), 62(10), 818-825 CODEN: JCLPDE; ISSN: 0160-6689
- PB Physicians Postgraduate Press, Inc.
- DT Journal
- LA English
- OSC.G 58 THERE ARE 58 CAPLUS RECORDS THAT CITE THIS RECORD (58 CITINGS)
- RE.CNT 50 THERE ARE 50 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 15 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Focus on ziprasidone
- AB A review. Ziprasidone is a new antipsychotic with combined dopamine and

serotonin receptor antagonist activity. The initial evidence suggests an effective dosage range of 80-160 mg/day. Clin. trials suggest that the drug is an effective antipsychotic in schizophrenia and schizoaffective disorder with a beneficial effect on neg. symptoms and symptoms of depression. The main adverse effects appear to be somnolence (14%) and nausea (10%). Ziprasidone has relatively fewer side-effects and yet has at least equivalent efficacy for florid "pos." symptoms compared with conventional antipsychotics. The addnl. serotonergic actions deliver further efficacy against "neg." and affective symptoms of schizophrenia. Reduced effects on cognitive abilities compared to conventional antipsychotics make ziprasidone more attractive still. Barring any unforeseen complications, it appears to a most valuable addition to the antipsychotic agents.

- AN 2001:903897 HCAPLUS <<LOGINID::20100601>>
- DN 136:177344
- TI Focus on ziprasidone
- AU Green, Ben
- CS Halton Hospital and the University of Liverpool, Liverpool, UK
- SO Current Medical Research and Opinion (2001), 17(2), 146-150 CODEN: CMROCX; ISSN: 0300-7995
- PB LibraPharm Ltd.
- DT Journal; General Review
- LA English
- OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)
- RE.CNT 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 16 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI In vivo 5-HT2A receptor blockade by quetiapine. An R91150 single photon emission tomography study
- AΒ Background: Atypical antipsychotic drugs are thought to show a high degree of 5-HT2A receptor blockade, which may prevent the emergence of extrapyramidal symptoms. Method: 5-HT2A binding was estimated using 123I-5-I-R91150 and single photon emission tomog. (SPET) in 6 schizophrenic subjects treated with quetiapine at a mean daily dose of 350 mg for at least 5 wk and a matched sample of 6 healthy volunteers. Clin. and side-effect ratings were performed at baseline and at the time of SPET scanning. The reference region approach was used to define a 5-HT2A binding index in the frontal and temporal cortex. Results: Quetiapine treatment resulted in a decline in 5-HT2A receptor availability in the frontal cortex (mean 0.98) relative to healthy volunteers (mean 1.33 ± 0.16). All patients showed improvements in clin. symptom or side-effect ratings. The mean frontal cortex:cerebellum ratio after quetiapine treatment was neg. correlated with reduction in the Abnormal Involuntary Rating scale and Simpson-Angus scores (Bonferroni corrected), but not with the reduction in the scores from the scale for the assessment of pos. symptoms, the scale for the assessment of neg. symptoms, the Montgomery-Asberg depression rating scale or patient age. Conclusion: Quetiapine treatment results in significant in vivo blockade of cortical 5-HT2A, similar to other atypical antipsychotic drugs. This effect may contribute to its placebo level extrapyramidal side-effect profile.
- AN 2001:743627 HCAPLUS <<LOGINID::20100601>>
- DN 136:63991
- ${\tt TI}$ In vivo 5-HT2A receptor blockade by quetiapine. An R91150 single photon emission tomography study
- AU Jones, Hugh M.; Travis, Michael J.; Mulligan, Rachel; Bressan, Rodrigo A.; Visvikis, Dmitri; Gacinovic, Sveto; Ell, Peter J.; Pilowsky, Lyn S.
- CS Department of Psychological Medicine, Section of Neurochemical Imaging, Institute of Psychiatry, London, SE5 8AF, UK
- SO Psychopharmacology (Berlin, Germany) (2001), 157(1), 60-66 CODEN: PSCHDL; ISSN: 0033-3158

PB Springer-Verlag

DT Journal

LA English

OSC.G 8 THERE ARE 8 CAPLUS RECORDS THAT CITE THIS RECORD (8 CITINGS)
RE.CNT 38 THERE ARE 38 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L15 ANSWER 17 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI A double-blind, randomized, prospective evaluation of the efficacy and safety of risperidone versus haloperidol in the treatment of schizoaffective disorder
- The relative efficacy and safety of risperidone vs. haloperidol in the AB treatment of schizo-affective disorder was studied. Sixty-two patients (29 depressed type; 33 bipolar type) entered a three-site, randomized, double-blind, 6-wk trial of risperidone (up to 10 mg/day) or haloperidol (up to 20 mg/day). Trained raters assessed baseline, weekly, and end-of-study levels of psychopathol. with the Pos. and Neg. Syndrome Scale (PANSS), the 24-item Hamilton Rating Scale for Depression (HAM-D-24) and the Clinician-Administered Rating Scale for Mania (CARS-M). The authors were unable to statistically distinguish between risperidone and haloperidol in the amelioration of psychotic and manic symptoms. In addition, there was no difference in worsening of mania between the two agents in either subgroup (i.e., depressed or bipolar subgroups). For the total PANSS, risperidone produced a mean decrease of 16 points from baseline compared with a 14-point decrease with haloperidol. For the total CARS-M scale, risperidone and haloperidol produced mean change scores of 5 and 8 points, resp., and for the CARS-M Mania subscale, 3 and 7 points, resp. Addnl., risperidone produced a mean decrease of 13 points from the baseline 24-item HAM-D, compared with an 8-point decrease with haloperidol. In those patients who had more severe depressive symptoms (i.e., HAM-D baseline score >20), risperidone produced at least a 50% mean improvement in 12 (75%) of 16 patients in comparison to 8 (38%) of 21 patients receiving haloperidol. Haloperidol produced significantly more extrapyramidal side effects and resulted in more dropouts caused by any side effect. There was no difference between risperidone and haloperidol in reducing both psychotic and manic symptoms in this group of patients with schizo-affective disorder. Risperidone did not demonstrate a propensity to precipitate mania and was better tolerated than haloperidol.

In those subjects with higher baseline HAM-D scores (i.e., >20), risperidone produced a greater improvement in depressive symptoms than haloperidol.

AN 2001:597245 HCAPLUS <<LOGINID::20100601>>

DN 135:339096

- TI A double-blind, randomized, prospective evaluation of the efficacy and safety of risperidone versus haloperidol in the treatment of schizoaffective disorder
- AU Janicak, Philip G.; Keck, Paul E., Jr.; Davis, John M.; Kasckow, John W.; Tugrul, Karen; Dowd, Sheila M.; Strong, Jane; Sharma, Rajiv P.; Strakowski, Stephen M.
- CS The Psychiatric Clinical Research Center and Department of Psychiatry, College of Medicine, University of Illinois at Chicago, Chicago, IL, USA
- SO Journal of Clinical Psychopharmacology (2001), 21(4), 360-368 CODEN: JCPYDR; ISSN: 0271-0749
- PB Lippincott Williams & Wilkins
- DT Journal
- LA English
- OSC.G 12 THERE ARE 12 CAPLUS RECORDS THAT CITE THIS RECORD (12 CITINGS)
- RE.CNT 35 THERE ARE 35 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L15 ANSWER 18 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Treatment of suicidality in schizophrenia
- A review with 48 refs. Between 4 and 13% of people with schizophrenia AB commit suicide and between 25 and 50% make a suicide attempt, a reflection of the devastating toll this syndrome takes on the quality of life, i.e., the subjective and objective sense of well-being. Many risk factors for suicide in schizophrenia have been identified, the most important of which are previous suicide attempts, depression, hopelessness, substance abuse, and male gender. Insight into having a serious mental illness and less severe cognitive impairment are also associated with increased risk for suicide in schizophrenia, most likely when accompanied by feelings of hopelessness. Typical neuroleptic drugs have not been shown to reduce the risk of suicide. However, several types of evidence suggest that clozapine, an atypical antipsychotic drug, appreciably reduces the suicide attempt and completion rates in schizophrenia and schizo-affective disorder, perhaps by as much as 75-85%. Other atypical antipsychotic drugs may have a similar effect, but direct evidence is lacking. Improvement in pos. and neg. symptoms, reduced extrapyramidal side effects (EPS), a direct antidepressant action, improved cognitive function, and improved compliance may contribute to reduced suicidality. The International Suicide Prevention Trial (InterSePT) is a large prospective, randomized study intended to compare the effectiveness of clozapine with that of olanzapine in reducing suicide and suicide-related events in schizophrenic and schizoaffective patients. Some information about suicidality in the patient sample is reported here.
- AN 2001:480353 HCAPLUS <<LOGINID::20100601>>
- DN 135:266558
- TI Treatment of suicidality in schizophrenia
- AU Meltzer, Herbert Y.
- CS Division of Psychopharmacology, Vanderbilt University School of Medicine, Nashville, TN, 37212, USA
- SO Annals of the New York Academy of Sciences (2001), 932(Clinical Science of Suicide Prevention), 44-60 CODEN: ANYAA9; ISSN: 0077-8923
- PB New York Academy of Sciences
- DT Journal; General Review
- LA English
- OSC.G 12 THERE ARE 12 CAPLUS RECORDS THAT CITE THIS RECORD (12 CITINGS)
- RE.CNT 48 THERE ARE 48 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 19 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI The combination of a serotonin reuptake inhibitor and a 5-HT2C antagonist, inverse agonist or partial agonist
- AB The present invention relates to the use of compds. and compns. of compds. having serotonin reuptake inhibiting activity and 5-HT2C antagonistic, partial agonistic or inverse agonistic activity for the for the treatment of depression and other affective disorders. The combined serotonin reuptake inhibiting effect and the 5-HT2C antagonistic, partial agonistic or inverse agonistic effect may reside within the same chemical compound or in two different chemical compds. E.g., simultaneous administration of 10 $\mu \text{mol/kg}$ citalopram with 1 $\mu \text{mol/kg}$ RS 102221 or Lu 27121 showed significant enhancement of the effect of citalopram in rats.
- AN 2001:434808 HCAPLUS <<LOGINID::20100601>>
- DN 135:41033
- TI The combination of a serotonin reuptake inhibitor and a 5-HT2C antagonist, inverse agonist or partial agonist
- IN Cremers, Thomas Ivo Franciscus Hubert; Wikstroem, Hakan Wilhelm; Den Boer, Johan Antonie; Bosker, Fokko Jan; Westerink, Bernard Hendrik Cornelis;

Bogeso, Klaus Peter; Hogg, Sandra; Mork, Arne

PA H. Lundbeck A/s, Den.
SO PCT Int. Appl., 29 pp.
CODEN: PIXXD2

Patent English DT

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    EP 2003-27672
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ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT
OSC.G 7 THERE ARE 7 CAPLUS RECORDS THAT CITE THIS RECORD (7 CITINGS)
RE.CNT 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L15 ANSWER 20 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Long-term olanzapine therapy in the treatment of bipolar I disorder: An open-label continuation phase study
- AΒ Olanzapine has demonstrated efficacy in the treatment of acute mania in 2 double-blind, placebo-controlled trials. We describe the results of the open-label extension from one of these trials. In a 3-wk, double-blind study of patients with DSM-IV bipolar 1 disorder, olanzapine was superior to placebo for the treatment of acute manic symptoms. Of the 139 patients who entered the double-blind phase of the 3-wk study, 113 patients continued into the 49-wk open-label extension. Efficacy measurements including the Young Mania Rating Scale (YMRS), the 21-item Hamilton Rating Scale for Depression (HAM-D-21), the Clin. Global Impressions scale-Bipolar Version, and the Pos. and Neg. Syndrome Scale and safety measurements including the Simpson-Angus scale, the Barnes Akathisia Scale, and the Abnormal Involuntary Movement Scale were completed throughout. The anal. considered all treatment results, starting with the first olanzapine dose. Adjunctive lithium and fluoxetine were allowed during the open-label extension. The mean length of olanzapine treatment was 6.6 mo, with a mean modal dose of 13.9 mg/day. A significant mean improvement in the YMRS total score, baseline to endpoint (-18.01, p <.001), was observed During treatment, 88.3% of patients experienced a remission of manic symptoms (YMRS total score \leq 12), and only 25.5% subsequently relapsed (YMRS total score ≤ 15). Significant improvement in HAM-D-21 scores was observed (p < .001). Forty-one percent of patients were maintained on olanzapine monotherapy. The most common treatment-emergent adverse events reported were somnolence (46.0%), depression (38.9%), and weight gain (36.3%). During up to 1 yr of olanzapine therapy, either as monotherapy or in combination with lithium and/or fluoxetine, patients with bipolar disorder demonstrated significant improvement in mania and depression symptoms with a favorable safety profile. Further double-blind, controlled studies are needed to confirm these results.
- AN 2001:424523 HCAPLUS <<LOGINID::20100601>>
- DN 135:267049
- TI Long-term olanzapine therapy in the treatment of bipolar I disorder: An open-label continuation phase study
- AU Sanger, Todd M.; Grundy, Starr L.; Gibson, P. Joseph; Namjoshi, Madhav A.; Greaney, Michael G.; Tohen, Mauricio F.
- CS Lilly Corporate Center, Lilly Research Laboratories, Eli Lilly and Company, Indianapolis, IN, USA
- SO Journal of Clinical Psychiatry (2001), 62(4), 273-281 CODEN: JCLPDE; ISSN: 0160-6689
- PB Physicians Postgraduate Press, Inc.
- DT Journal
- LA English

- OSC.G 53 THERE ARE 53 CAPLUS RECORDS THAT CITE THIS RECORD (53 CITINGS)
- RE.CNT 43 THERE ARE 43 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 21 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI First experiences in combination therapy using olanzapine with SSRIs (citalopram, paroxetine) in delusional depression
- AB In an open prospective study, 26 patients with delusional depression (mood-congruent psychotic features: DSMIV 296.4) were treated over 5 wk with a combination of SSRI (citalopram, n = 22, or paroxetine, n = 4) and the neuroleptic olanzapine. The course of therapy was evaluated with the Hamilton depression scale (HAMD). Not only the total HAMD score, but also the subscores for affectivity and delusional symptoms decreased significantly. After the end of the 5-wk combination therapy, 18 out of 26 patients (69%) could be discharged as responders to outpatient treatment. The course of treatment was characterized by excellent tolerance.
- AN 2001:384296 HCAPLUS <<LOGINID::20100601>>
- DN 135:236300
- TI First experiences in combination therapy using olanzapine with SSRIs (citalopram, paroxetine) in delusional depression
- AU Konig, F.; Hippel, C. V.; Petersdorff, T.; Neuhoffer-Weiss, M.; Wolfersdorf, M.; Kaschka, W. P.
- CS Department of Psychiatry 1, University of Ulm, Ravensburg, Germany
- SO Neuropsychobiology (2001), 43(3), 170-174 CODEN: NPBYAL; ISSN: 0302-282X
- PB S. Karger AG
- DT Journal
- LA English
- OSC.G 10 THERE ARE 10 CAPLUS RECORDS THAT CITE THIS RECORD (10 CITINGS)
- RE.CNT 30 THERE ARE 30 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 22 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Combination treatment for depression and anxiety containing a CNS-penetrant NK-1 receptor antagonist and an antidepressant or anxiolytic agent
- AB A combination treatment for depression and anxiety contains a CNS-penetrant NK-1 receptor antagonist and an antidepressant or anxiolytic agent. Tablets were prepared containing a NK-1 antagonist and sertraline or ziprasidone.
- AN 2001:356210 HCAPLUS <<LOGINID::20100601>>
- DN 134:357580
- TI Combination treatment for depression and anxiety containing a CNS-penetrant NK-1 receptor antagonist and an antidepressant or anxiolytic agent
- IN Sobolov-Jaynes, Susan Beth
- PA Pfizer Products Inc., USA
- SO Eur. Pat. Appl., 46 pp. CODEN: EPXXDW
- DT Patent
- LA English
- FAN.CNT 1

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OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)
RE.CNT 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L15 ANSWER 23 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Novel, highly potent and selective serotonin 5-HT2A/dopamine D2 receptor antagonists as potential antipsychotics
- The 5-HT2A and D2 receptors have been implicated as therapeutic targets AΒ for schizophrenia and depression as well as other neuropsychiatric diseases. The atypical antipsychotic agents possess dual 5-HT2A/D2 antagonism and have demonstrated superior clin. efficacy for schizophrenia with a reduced propensity to induce extrapyramidal side effect compared to typical antipsychotic agents (dopamine D2 receptor antagonists). However, identification of ligands with proper 5-HT2A/D2 receptor binding ratios and selectivities >100-fold vs. other monoamine receptors and the various neurotransmitter transporters has not been achieved. The most widely investigated atypical antipsychotics, clozapine and risperidone, are the standard for this class of agents exhibiting potent 5-HT2A antagonism, moderate D2 affinity and only modest selectivity over a wide range of receptors. We will describe our recent efforts in the area of selective dual 5-HT2A/D2 antagonists for potential use as atypical antipsychotics. The strategy for tailoring 5-HT2A/D2 receptor-binding affinity ratios will also be discussed. Structure activity studies of a novel series of compds. have led to the identification of orally bioavailable, highly potent and selective ligands for the target receptor subtypes that demonstrate efficacy in rat behavioral models for 5-HT2A and D2 antagonism.
- AN 2001:201996 HCAPLUS <<LOGINID::20100601>>
- TI Novel, highly potent and selective serotonin 5-HT2A/dopamine D2 receptor antagonists as potential antipsychotics
- AU Lee, Taekyu; Robichaud, Albert J.; Boyle, Kristopher E.; Lu, Yimin; Chen, Wenting; McClung, Christopher; Deng, Wei; Miller, Keith J.; McElroy, John F.; Largent, Brian L.
- CS Department of Medicinal Chemistry, DuPont Pharmaceuticals Company, Wilmington, DE, 19880-0500, USA
- SO Abstracts of Papers, 221st ACS National Meeting, San Diego, CA, United States, April 1-5, 2001 (2001) MEDI-098 CODEN: 69FZD4
- PB American Chemical Society
- DT Journal; Meeting Abstract
- LA English
- L15 ANSWER 24 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Olanzapine: An updated review of its use in the management of schizophrenia
- AB A review with 307 refs. Olanzapine, a thienobenzodiazepine derivative, is a second generation (atypical) antipsychotic agent which has proven efficacy against the pos. and neg. symptoms of schizophrenia. Compared with conventional antipsychotics, it has greater affinity for serotonin 5-HT2A than for dopamine D2 receptors. In large, well controlled trials in patients with schizophrenia or related psychoses, olanzapine 5 to 20 mg/day was significantly superior to haloperidol 5 to

20 mg/day in overall improvements in psychopathol. rating scales and in the treatment of depressive and neg. symptoms, and was comparable in effects on pos. psychotic symptoms. The 1-yr risk of relapse (rehospitalization) was significantly lower with olanzapine than with haloperidol treatment. In the first double-blind comparative study (28-wk) of olanzapine and risperidone, olanzapine 10 to 20 mg/day proved to be significantly more effective than risperidone 4 to 12 mg/day in the treatment of neq. and depressive symptoms but not on overall psychopathol. symptoms. In contrast, preliminary results from an 8-wk controlled study suggested risperidone 2 to 6 mg/day was superior to olanzapine 5 to 20 mg/day against pos. and anxiety/depressive symptoms (p < 0.05), although consistent with the first study, both agents demonstrated similar efficacy on measures of overall psychopathol. Improvements in general cognitive function seen with olanzapine treatment in a 1-yr controlled study of patients with early-phase schizophrenia, were significantly greater than changes seen with either risperidone or haloperidol. However, preliminary results from an 8-wk trial showed comparable cognitive enhancing effects of olanzapine and risperidone treatment in patients with schizophrenia or schizo-affective disorder. Several studies indicate that olanzapine has benefits against symptoms of aggression and agitation, while other studies strongly support the effectiveness of olanzapine in the treatment of depressive symptomatol. Olanzapine is associated with significantly fewer extrapyramidal symptoms than haloperidol and risperidone. In addition, olanzapine is not associated with a risk of agranulocytosis as seen with clozapine or clin. significant hyperprolactinemia as seen with risperidone or prolongation of the QT interval. The most common adverse effects reported with olanzapine are bodyweight gain, somnolence, dizziness, anticholinergic effects (constipation and dry mouth) and transient asymptomatic liver enzyme elevations. In comparison with haloperidol, the adverse events reported significantly more frequently with olanzapine in ≥3.5% of patients were dry mouth, bodyweight gain and increased appetite and compared with risperidone, only bodyweight gain occurred significantly more frequently with olanzapine. The high acquisition cost of olanzapine is offset by redns. in other treatment costs (inpatient and/or outpatient services) of schizophrenia. Pharmacoeconomic analyses indicate that olanzapine does not significantly increase, and may even decrease, the overall direct treatment costs of schizophrenia, compared with haloperidol. Compared with risperidone, olanzapine has also been reported to decrease overall treatment costs, despite the several-fold higher daily acquisition cost of the drug. Olanzapine treatment improves quality of life in patients with schizophrenia and related psychoses to a greater extent than haloperidol, and to broadly the same extent as risperidone. Conclusions: Olanzapine demonstrated superior antipsychotic efficacy compared with haloperidol in the treatment of acute phase schizophrenia, and in the treatment of some patients with first-episode or treatment-resistant schizophrenia. The reduced risk of adverse events and therapeutic superiority compared with haloperidol and risperidone in the treatment of neg. and depressive symptoms support the choice of olanzapine as a first-line option in the management of schizophrenia in the acute phase and for the maintenance of treatment response.

- AN 2001:141840 HCAPLUS <<LOGINID::20100601>>
- DN 135:161855
- TI Olanzapine: An updated review of its use in the management of schizophrenia
- AU Bhana, Nila; Foster, Rachel H.; Olney, Roger; Plosker, Greg L.
- CS Adis International Limited, Auckland, N. Z.
- SO Drugs (2001), 61(1), 111-161 CODEN: DRUGAY; ISSN: 0012-6667
- PB Adis International Ltd.
- DT Journal; General Review

LA English

OSC.G 64 THERE ARE 64 CAPLUS RECORDS THAT CITE THIS RECORD (64 CITINGS) RE.CNT 309 THERE ARE 309 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L15 ANSWER 25 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN

- TI Ziprasidone in the short-term treatment of patients with schizoaffective disorder: results from two double-blind, placebo-controlled, multicenter studies
- This study assessed the efficacy of ziprasidone for the treatment of AB schizoaffective disorder. Data were taken from subsets of patients with schizoaffective disorder, derived from two sep. double-blind, placebo-controlled, parallel-group, multicenter studies. A total of 115 hospitalized patients with an acute episode of schizoaffective disorder were randomly assigned to receive either fixed oral doses of ziprasidone 40 mg/day (N = 16), 80 mg/day (N = 18), 120 mg/day (N = 22), 160 mg/day (N = 25), or placebo (N = 34) for 4 to 6 wk. Mean baseline-to-endpoint changes in Brief Psychiatric Rating Scale (BPRS) total, BPRS Core, Clin. Global Impressions Severity scale (CGI-S), BPRS Depressive, BPRS Manic, and Montgomery-Åsberg Depression Rating Scale total scores were compared between the placebo and ziprasidone groups. Neurol. (Simpson-Angus, Barnes Akathisia, Abnormal Involuntary Movement Scale [AIMS]) and other side effects were also assessed. Significant dose-related improvements on all primary efficacy variables (BPRS total, BPRS Core, CGI-S) and for BPRS Manic items were observed with ziprasidone treatment in a combined anal. of data from both studies ($p \le 0.01$). Ziprasidone 160 mg/day was significantly more effective than placebo in improving mean BPRS total, BPRS Core, BPRS Manic, and CGI-S scores (p < 0.05). At 120 mg/day, ziprasidone was significantly more effective than placebo in improving mean CGI-S scores (p < 0.05). The incidence of individual adverse events was generally low in all treatment groups and was not dose-related. In addition, no significant differences were observed between baseline-to-endpoint mean changes in Simpson-Angus and AIMS scores with placebo or ziprasidone 40 to 160 mg/day. These results suggest that ziprasidone may have efficacy in the treatment of affective as well as psychotic symptoms of schizoaffective disorder, with a low side-effect burden.
- AN 2001:93669 HCAPLUS <<LOGINID::20100601>>
- DN 135:132230
- TI Ziprasidone in the short-term treatment of patients with schizoaffective disorder: results from two double-blind, placebo-controlled, multicenter studies
- AU Keck, Paul E., Jr.; Reeves, Karen R.; Harrigan, Edmund P.
- CS Biological Psychiatry Program, University of Cincinnati College of Medicine, Cincinnati, OH, USA
- SO Journal of Clinical Psychopharmacology (2001), 21(1), 27-35 CODEN: JCPYDR; ISSN: 0271-0749
- PB Lippincott Williams & Wilkins
- DT Journal
- LA English
- OSC.G 34 THERE ARE 34 CAPLUS RECORDS THAT CITE THIS RECORD (34 CITINGS)
- RE.CNT 47 THERE ARE 47 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 26 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Olanzapine: Preclinical and clinical profiles of a novel antipsychotic agent
- AB A review with 278 refs. The novel antipsychotic agent olanzapine (Zyprexa, Eli Lilly and Company) is a thienobenzodiazepine analog marketed for the treatment of schizophrenia. Olanzapine's diverse receptor binding profile and greater affinity for serotonin receptors over

dopamine receptors is thought to impart antipsychotic efficacy with a low incidence of serious extrapyramidal symptoms (EPS). With once daily dosing steady-state plasma concns. reached within approx. 1 wk. Olanzapine is extensively metabolized by the liver, is mostly excreted in the urine, and has few drug intractions. In clin. trials, the efficacy of olanzapine for treating schizophrenia is better than placebo and haloperidol and comparable to risperidone. Olanzapine may also ameliorate some comorbid symptoms including neg. symptoms, depression, anxiety, substance abuse, and cognitive dysfunction, and it is effective in the long-term maintenance of response, treatment-resistance, and improving quality of life. The overall direct costs are lower with olanzapine treatment compared with haloperidol or risperidone treatment. In clin. trials, olanzapine demonstrates a favorable safety profile. The most frequently reported treatment-emergent adverse events are somnolence, schizophrenic reaction, insomnia, headache, agitation, rhinitis, and weight gain. Significantly fewer EPS (based on formal rating scales) and incidences of tardive dyskinesia have been reported for olanzapine compared with haloperidol. Olanzapine has not been associated with persistent elevations of prolactin above the upper limit of normal nor has it been associated with clin. significant changes in cardiac QTc interval. Fewer incidences of suicide attempts have been reported with olanzapine compared with placebo, haloperidol, or risperidone treatments. There is evidence that olanzapine may be effective in the treatment of mood disorders, psychosis associated with. Alzheimer's disease, obsessive-compulsive disorder, pervasive developmental disorders, and delirium. Patients with schizophrenia have been successfully switched from other antipsychotics to olanzapine. In conclusion, olanzapine offers a significantly improved risk-to-benefit profile compared with haloperidol and possibly risperidone, and thus should be considered an important treatment option for schizophrenia and related disorders.

- AN 2001:49031 HCAPLUS <<LOGINID::20100601>>
- DN 135:86379
- TI Olanzapine: Preclinical and clinical profiles of a novel antipsychotic agent
- AU Tollefson, Gary D.; Taylor, Cindy C.
- CS Lilly Research Laboratories Lilly Corporate Center, Eli Lilly and Company, Indianapolis, IN, 46285, USA
- SO CNS Drug Reviews (2000), 6(4), 303-363 CODEN: CDREFB; ISSN: 1080-563X
- PB Neva Press
- DT Journal; General Review
- LA English
- OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)
- RE.CNT 278 THERE ARE 278 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 27 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Methods and compositions for the treatment of neuroleptic and related disorders using sertindole derivatives
- AB The invention relates to novel methods using, and pharmaceutical compns. and dosage forms comprising, sertindole derivs. Sertindole derivs. include, but are not limited to, nor-sertindole, 5-oxo-sertindole, dehydro-sertindole, and dehydro-nor-sertindole. The methods of the invention are directed to the treatment and prevention of neuroleptic and related disorders such as, but are not limited to, psychotic disorders, depression, anxiety, substance addiction, memory impairment and pain. For example, capsules were prepared containing a sertindole derivative
- 50.0
- mg, lactose 48.5 mg, TiO2 0.5 mg, and Mg stearate 1.0 mg.
- AN 2000:861482 HCAPLUS <<LOGINID::20100601>>
- DN 134:32977

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disorders using sertindole derivatives
ΙN
     Jerussi, Thomas P.
     Sepracor Inc., USA
PA
     PCT Int. Appl., 33 pp.
SO
     CODEN: PIXXD2
DT
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     English
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L15 ANSWER 28 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
     Serotoninergic agonists and antagonists for treatment of
TΙ
     bronchoconstriction
     The present invention relates to a compound having agonist activity to the
AΒ
     5-HT4 receptor or antagonist activity to the 5-HT2a receptor and manufacture of
     a medicament for prophylactic or therapeutic treatment of disorders
     involving bronchoconstriction of a human or animal, such as asthma,
     emphysema, chronic bronchitis, chronic obstructive pulmonary disease,
     depression, anorectic or bulimic eating disorders, anxiety or
     various psychotic conditions including schizophrenia. Compds. of the
     present invention have the capacity of reducing the pathol.
     bronchoconstriction by at least 30%, preferably at least 60%, and most
     preferably at least 90%.
ΑN
    2000:772451 HCAPLUS <<LOGINID::20100601>>
    133:329581
DΝ
ΤI
     Serotoninergic agonists and antagonists for treatment of
     bronchoconstriction
     Skogvall, Staffan
IN
     Respiratorius AB, Swed.
PΑ
     PCT Int. Appl., 28 pp.
SO
     CODEN: PIXXD2
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LA
    English
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    WO 2000064441 A2 20001102
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Methods and compositions for the treatment of neuroleptic and related

ТΤ

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         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
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- L15 ANSWER 29 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Nefazodone in the adjunctive therapy of schizophrenia: An open-label exploratory study
- AΒ Compared to conventional antipsychotic medications, atypical antipsychotic medications demonstrate greater central serotonin (5HT2) receptor antagonism than dopamine type 2 (D2) receptor antagonism. Nefazodone, an antidepressant medication, exhibits 5HT2 receptor antagonism; we therefore wondered if its addition to stable regimens of antipsychotic medication would increase antipsychotic efficacy, independently of a primary effect on mood, through the mechanism of augmented 5HT2 receptor antagonism. In a pilot investigation, we administered nefazodone (400 mg/d) for 6 wk as an open-label adjunct to antipsychotic medication in 10 patients with chronic schizophrenia. The patients were moderately depressed at baseline but did not meet criteria for major depressive episode. The Brief Psychiatric Rating Scale (BPRS) and Montgomery-Asberg Depression Rating Scale scores showed statistically significant and clin. robust improvements with nefazodone treatment, which were maintained at follow-up evaluation 2 wk after the end of nefazodone treatment. There were no adverse events. These results suggest that nefazodone may be a safe and effective adjunct to antipsychotic medications in schizophrenia and that augmentation of

5HT2 antagonism may prove to be a viable strategy for "boosting" antipsychotic efficacy and for treating depressive symptoms in schizophrenia.

- AN 2000:753368 HCAPLUS <<LOGINID::20100601>>
- DN 134:320731
- TI Nefazodone in the adjunctive therapy of schizophrenia: An open-label exploratory study
- AU Rosenberg, Paul B.; Rosse, Richard B.; Schwartz, Barbara L.; Deutsch, Stephen I.
- CS Mental Health Service Line, Georgetown University School of Medicine, Washington, DC, USA
- SO Clinical Neuropharmacology (2000), 23(4), 222-225 CODEN: CLNEDB; ISSN: 0362-5664
- PB Lippincott Williams & Wilkins
- DT Journal
- LA English
- OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)
- RE.CNT 28 THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 30 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI A novel approach to the identification of psychiatric drugs: serotonin-glutamate interactions in the prefrontal cortex
- AB A review with 83 refs. Activation of neocortical 5-hydroxytryptamine2A (5-HT2A) receptors is thought to mediate the profound psychomimetic effects of hallucinogenic drugs such as LSD and mescaline. These effects include alteration in mood, perception, and cognition. Conversely, blockade of neocortical 5-HT2A receptor may be related to the thymoleptic effects of newly released antidepressant (e.g., mirtazepine, nefazodone) and atypical antipsychotic drugs (e.g., risperidone, olanzapine). Therefore, one strategy to develop novel antidepressant drugs might be to identify drugs which suppress the effects of 5-HT2A receptor activation in key neurocircuits. Electrophysiol. expts. using in vitro rat slices of the medial prefrontal cortex have found that activation of 5-HT2A receptors results in glutamate release from thalamocortical terminals by a novel focal effect. A number of monoamine (5-HT1/7, β 2), metabotropic glutamate (mGlu2), and neuropeptide (μ -opioid) receptors suppress the glutamate release induced by 5-HT2A receptor activation. Clin. studies examining the effects of serotonin or catecholamine depletion suggest the activation of 5-HT or catecholamine receptors mediate the therapeutic effects of selective serotonin reuptake inhibitors (SSRIs) or tricyclic antidepressants (TCAs), resp. In addition, opiate agonists may have antidepressant properties. Therefore, it is suggested that elucidation of the specific receptors that suppress glutamate release induced by 5-HT2A receptor activation in the medial prefrontal cortex may have several effects. First, this might lead to a more complete understanding of the 5-HT receptor(s) that mediate the therapeutic effects of presently used drugs such as SSRIs. This site might be a therapeutic target free of side effects such as sexual dysfunction. Second, this strategy might lead to novel therapeutic targets for depression, such as metabotropic glutamate agonists which may not be efficacious in screening strategies primarily dependent on synaptic availability of monoaminergic neurotransmitters.
- AN 2000:716745 HCAPLUS <<LOGINID::20100601>>
- DN 134:25436
- TI A novel approach to the identification of psychiatric drugs: serotonin-glutamate interactions in the prefrontal cortex
- AU Marek, Gerard J.
- CS Yale School of Medicine, New Haven, CT, 06508, USA
- SO CNS Drug Reviews (2000), 6(3), 206-218

CODEN: CDREFB; ISSN: 1080-563X

PB Neva Press

DT Journal; General Review

LA English

OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS) RE.CNT 83 THERE ARE 83 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L15 ANSWER 31 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI A double-blind, placebo-controlled study of risperidone addition in serotonin reuptake inhibitor-refractory obsessive-compulsive disorder
- To date, only 1 controlled study has found a drug (haloperidol) to be AΒ efficacious in augmenting response in patients with obsessive-compulsive disorder (OCD) refractory to serotonin reuptake inhibitor (SRI) monotherapy; patients with comorbid chronic tic disorders showed a preferential response. This report describes the first controlled study of risperidone addition in patients with OCD refractory to treatment with SRI alone. Seventy adult patients with a primary DSM-IV diagnosis of OCD received 12 wk of treatment with an SRI. Thirty-six patients were refractory to the SRI and were randomized in a double-blind manner to 6 wk of risperidone (n=20) or placebo (n=16) addition Behavioral ratings, including the Yale-Brown Obsessive Compulsive Scale, were obtained at baseline and throughout the trial. Placebo-treated patients subsequently received an identical open-label trial of risperidone addition For study completers, 9 (50%) of 18 risperidone-treated patients were responders (mean daily dose, $2.2\pm0.7~\text{mg/d}$) compared with 0 of 15 in the placebo addition group (P<.005). Seven (50%) of 14 patients who received open-label risperidone addition responded. Risperidone addition was superior to placebo in
- reducing OCD (P<.001), depressive (P<.001), and anxiety (P=.003) symptoms. There was no difference in response between OCD patients with and without comorbid diagnoses of chronic tic disorder or schizo-typal personality disorder. Other than mild, transient sedation, risperidone was well tolerated. These results suggest that OCD patients with and without comorbid chronic tic disorders or schizo-typal personality disorder may respond to the addition of low-dose risperidone to ongoing SRI therapy.
- AN 2000:596352 HCAPLUS <<LOGINID::20100601>>
- DN 134:80743
- TI A double-blind, placebo-controlled study of risperidone addition in serotonin reuptake inhibitor-refractory obsessive-compulsive disorder
- AU McDougle, Christopher J.; Epperson, C. Neill; Pelton, Gregory H.; Wasylink, Suzanne; Price, Lawrence H.
- CS Department of Psychiatry, Indiana University School of Medicine, Indianapolis, USA
- SO Archives of General Psychiatry (2000), 57(8), 794-801 CODEN: ARGPAQ; ISSN: 0003-990X
- PB American Medical Association
- DT Journal
- LA English
- OSC.G 108 THERE ARE 108 CAPLUS RECORDS THAT CITE THIS RECORD (108 CITINGS)
- RE.CNT 31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 32 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Synergistic Effects of Olanzapine and Other Antipsychotic Agents in Combination with Fluoxetine on Norepinephrine and Dopamine Release in Rat Prefrontal Cortex
- AB To understand the mechanism of the clin. efficacy of olanzapine and

fluoxetine combination therapy for treatment-resistant depression (TRD), we studied the effects of olanzapine and other antipsychotics in combination with the selective serotonin uptake inhibitors fluoxetine or sertraline on neurotransmitter release in rat prefrontal cortex (PFC) using microdialysis. The combination of olanzapine and fluoxetine produced robust, sustained increases of extracellular levels of dopamine ([DA]ex) and norepinephrine ([NE]ex) up to 361±28% and 272±16% of the baseline, resp., which were significantly greater than either drug alone. This combination produced a slightly smaller increase of serotonin ([5-HT]ex) than fluoxetine alone. The combination of clozapine or risperidone with fluoxetine produced less robust and persistent increases of [DA]ex and [NE]ex. The combination of haloperidol or MDL 100907 with fluoxetine did not increase the monoamines more than fluoxetine alone. Olanzapine plus sertraline combination increased only [DA]ex. Therefore, the large, sustained increase of [DA]ex, [NE]ex, and [5-HT]ex in PFC after olanzapine-fluoxetine treatment was unique and may contribute to the profound antidepressive effect of the olanzapine and fluoxetine therapy in TRD.

- AN 2000:565105 HCAPLUS <<LOGINID::20100601>>
- DN 134:125804
- TI Synergistic Effects of Olanzapine and Other Antipsychotic Agents in Combination with Fluoxetine on Norepinephrine and Dopamine Release in Rat Prefrontal Cortex
- AU Zhang, W.; Perry, K. W.; Wong, D. T.; Potts, B. D.; Bao, J.; Tollefson, G. D.; Bymaster, F. P.
- CS Neuroscience Research Division, Lilly Research Laboratories, Lilly Corporate Center, Indianapolis, IN, USA
- SO Neuropsychopharmacology (2000), 23(3), 250-262 CODEN: NEROEW; ISSN: 0893-133X
- PB Elsevier Science Inc.
- DT Journal
- LA English
- OSC.G 105 THERE ARE 105 CAPLUS RECORDS THAT CITE THIS RECORD (106 CITINGS)
 RE.CNT 46 THERE ARE 46 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 33 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- ${\tt TI}$ Association analysis of the 5-HT5A gene in depression, psychosis and antipsychotic response
- AΒ The serotonergic system is targeted by both antidepressants and atypical antipsychotic drugs such as clozapine. Genetic variation in the 5-HT5A gene might be involved in susceptibility to depression or the major psychoses or in influencing clin. response to treatment. To examine this hypothesis, two polymorphisms (-19G/C; 12A/T) in the human 5-HT5A receptor gene were genotyped in a sample of 269 unrelated schizophrenic patients treated with clozapine, 112 bipolar patients, 75 unipolar patients, and 187 controls. After 5-fold correction for multiple testing, allelic association was found with the -19G/Cpolymorphism and bipolar affective disorder, unipolar depression , and schizophrenia, indicating a potential protective effect of the G19 allele. For the 12A/T polymorphism allelic association was observed with unipolar depression only. Thus, allelic variation in the human 5-HT5A receptor gene may be involved in susceptibility to schizophrenia and affective disorders but not in determining response to clozapine.
- AN 2000:478478 HCAPLUS <<LOGINID::20100601>>
- DN 133:348602
- TI Association analysis of the 5-HT5A gene in depression, psychosis and antipsychotic response
- AU Birkett, Joseph T.; Arranz, Maria J.; Munro, Janet; Osbourn, Sarah; Kerwin, Robert W.; Collier, David A.
- CS Section of Clinical Neuropharmacology, Division of Psychological Medicine,

Institute of Psychiatry, London, SE5 8AF, UK NeuroReport (2000), 11(9), 2017-2020 SO CODEN: NERPEZ; ISSN: 0959-4965 РΒ Lippincott Williams & Wilkins DΤ Journal English LA THERE ARE 42 CAPLUS RECORDS THAT CITE THIS RECORD (43 CITINGS) OSC.G 42 RE.CNT 28 THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 34 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN

Effects of the CRF1 receptor antagonist, CP 154,526, in the separation-induced vocalization anxiolytic test in rat pups AB CRF1 receptor antagonists have been proposed as novel pharmacol. treatments for depression, anxiety and stress disorders. The primary goal of the present study was to evaluate the effects of the CRF1 receptor antagonist, CP 154,526, in the separation-induced vocalization (SIV) model of anxiety. Nine- to 11-day-old rat pups were separated from their litter and the effects of i.p. administered test compds. on the elicited ultrasonic vocalizations were measured. Side-effect potential was assessed using a modified inclined plane test ('time on an inclined plane', or TIP), and using neg. geotaxis. SIV was reduced by CP 154,526 at doses that did not affect TIP or neq. geotaxis, a profile like that of the 5-HT1A partial agonist buspirone. The benzodiazepine anxiolytic, diazepam, decreased SIV but also produced significant side effects at one to three-fold higher doses. Addnl. pharmacol. characterization of SIV demonstrated anxiolytic-like effects of the atypical antipsychotic, clozapine, but not the typical antipsychotic, haloperidol, and of the serotonin reuptake inhibitor, zimelidine, but not the norepinephrine reuptake inhibitor, desipramine. In summary, the data support the conclusion that selective CRF1 receptor antagonists may have utility in anxiety and stress disorders. The data further support the use of separation-induced vocalizations for identifying mechanistically diverse compds. with anxiolytic actions in man.

AN 2000:322632 HCAPLUS <<LOGINID::20100601>>

DN 133:129772

L15

TI Effects of the CRF1 receptor antagonist, CP 154,526, in the separation-induced vocalization anxiolytic test in rat pups

AU Kehne, J. H.; Coverdale, S.; McCloskey, T. C.; Hoffman, D. C.; Cassella, J. V.

CS Neurogen Corporation, Branford, CT, 06405, USA

SO Neuropharmacology (2000), 39(8), 1357-1367 CODEN: NEPHBW; ISSN: 0028-3908

PB Elsevier Science Ltd.

DT Journal

LA English

OSC.G 38 THERE ARE 38 CAPLUS RECORDS THAT CITE THIS RECORD (38 CITINGS)

RE.CNT 59 THERE ARE 59 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L15 ANSWER 35 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Atypical antipsychotic agents: a critical review
- AB A review with 155 refs. The pharmacol, efficacy, and adverse effects of atypical antipsychotic agents when used to treat schizophrenia and other disorders are reviewed. Atypical antipsychotic agents were developed in response to problems with typical agents, including lack of efficacy in some patients, lack of improvement in neg. symptoms, and troublesome adverse effects, especially extrapyramidal symptoms (EPSs) and tardive dyskinesia (TD). Atypical antipsychotics differ from typical psychotics in their "limbic-specific" dopamine type 2 (D2)-receptor binding and high ratio of serotonin

type 2 (5-HT2)-receptor binding to D2 binding. In clin. trials in patients with non-treatment-resistant schizophrenia, risperidone and olanzapine were superior to placebo for pos. and neg. symptoms. Risperidone and olanzapine were superior to haloperidol on some measures. Quetiapine was better than placebo but was not better than typical antipsychotics. Head-to-head comparisons of atypical antipsychotics in non-treatment-resistant schizophrenia have been inconclusive. Clozapine remains the standard agent for treatment-resistant schizophrenia. Atypical agents are substantially more expensive than their typical antipsychotic counterparts. To fully determine the overall efficiency of these drugs, other potential benefits, such as improved quality of life, need to be factored in. Atypical antipsychotics are associated with a decreased capacity to cause EPSs, TD, neuroleptic malignant syndrome, and hyperprolactinemia. Clozapine carries a risk of agranulocytosis; the white blood cell count must be monitored. Atypical antipsychotics are increasingly being used for indications other than schizophrenia, such as the management of aggression, mania, and depression. Atypical antipsychotics are often considered first-line agents for treating schizophrenia and are promising treatment alternatives for other psychiatric and neurol. conditions.

- AN 2000:110433 HCAPLUS <<LOGINID::20100601>>
- DN 132:146053
- TI Atypical antipsychotic agents: a critical review
- AU Worrel, Jodi A.; Marken, Patricia A.; Beckman, Stephanie E.; Ruether, Valerie L.
- CS Psychiatry, St. Cloud Veterans Affairs Medical Center, St. Cloud, MN, USA
- SO American Journal of Health-System Pharmacy (2000), 57(3), 238-255
 - CODEN: AHSPEK; ISSN: 1079-2082
- PB American Society of Health-System Pharmacists
- DT Journal; General Review
- LA English
- OSC.G 61 THERE ARE 61 CAPLUS RECORDS THAT CITE THIS RECORD (61 CITINGS)
- RE.CNT 155 THERE ARE 155 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 36 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Behavioral pharmacology of cis-flupentixol compared to typical and atypical neuroleptics, anxiolytics, and antidepressives
- AB The behavioral profile was studied of cis-flupentixol in various animal models in comparison with the typical neuroleptic haloperidol, the atypical neuroleptic clozapine, the benzodiazepinanxiolytic diazepam, and the tricyclic antidepressant amitriptyline. Flupentixol was effective in 2 psychosis and 1 anxiety model. Its profile showed similarities with that of clozapin and was contrary to G2that of haloperidol. Some similarities were found between flupentixol and amitriptylin in a depression model.
- AN 2000:250 HCAPLUS <<LOGINID::20100601>>
- DN 132:44886
- TI Behavioral pharmacology of cis-flupentixol compared to typical and atypical neuroleptics, anxiolytics, and antidepressives
- AU De Vry, J.
- CS Germany
- SO Flupentixol Typisches oder Atypisches Wirkspektrum?: Pharmakologie, Antipsychotische Wirkung, neue Indikationen (1998), 23-34.

 Editor(s): Glaser, T.; Soyka, M. Publisher: Dr. Dietrich Steinkopff Verlag GmbH & Co. KG, Darmstadt, Germany.

 CODEN: 68MEAY
- DT Conference
- LA German
- OSC.G 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)

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L15 ANSWER 37 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
     Combination therapy of atypical antipsychotics and serotonin
TΤ
     reuptake inhibitors for treatment of bipolar disorders
AB
     The invention provides methods and compns. for the treatment of bipolar
     disorder, bipolar depression or unipolar depression,
     all with or without psychotic features. This method employs a compound
     having activity as an atypical antipsychotic in
     combination with an effective amount of a second compound selected from the
     group consisting of a serotonin reuptake inhibitor, an
     anticonvulsant and lithium. Pharmaceutical formulations of combination of
     drugs of the invention are presented. E.g., hard gelatin capsules were
     prepared containing olanzapine 25 mg, fluoxetine-HCl 20 mg, starch 150 mg, and
     Mg stearate 10 mg. In a double blind trial in patients diagnosed with
     treatment-resistant major depression, the administration of
     fluoxetine plus olanzapine (20-60 mg/day and 5-20 mg/day, resp.) resulted
     in a greater improvement on the HAMD-21 score than either of the
     monotherapy.
     1999:783941 HCAPLUS <<LOGINID::20100601>>
ΑN
DN
     132:9033
ΤI
     Combination therapy of atypical antipsychotics and serotonin
     reuptake inhibitors for treatment of bipolar disorders
     Tollefson, Gary Dennis
ΙN
     Eli Lilly and Company, USA
PΑ
     PCT Int. Appl., 37 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 1
     PATENT NO.
                   KIND DATE APPLICATION NO. DATE
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         RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, BF, BJ, CF, CG, CI, CM,
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NZ 507981 A 20031031 NZ
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HR 2000000798 A2 20011031 HF
NO 2000005884 A 20010124 NC
ZA 2000006817 A 20020221 ZF
US 20030027817 A1 20030206 US
PRAI US 1998-87126P P 19980529 <--
WO 1999-US11314 W 19990521 <--
US 2000-700446 B1 20001109 <--
                                              MX 2000-11354
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                                                                      20001120 <--
                                              HR 2000-798
                                             NO 2000-5884 20001121 <--

ZA 2000-6817 20001121 <--

US 2002-165850 20020607 <--
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OSC.G 12 THERE ARE 12 CAPLUS RECORDS THAT CITE THIS RECORD (12 CITINGS)

- L15 ANSWER 38 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- ${\sf TI}$ Compositions and methods employing R(-)fluoxetine and other active ingredients
- AB Pharmaceutical compns. which comprise R(-) fluoxetine and one or more other biol. active compds. e.g. a benzodiazepine compound, a tricyclic antidepressant, a 5-HT1A receptor antagonist, a 5-HT3 receptor agonist, a β -adrenergic antagonist, an antipsychotic agent, an anti-anxiolytic or other psychotropic drug, are disclosed. Methods of treating or preventing a disease or disorder, especially a psychotic or psychiatric disease or disorder, using the above pharmaceutical composition or by administering a R(-)fluoxetine in combination with one or more other biol. active compds. are also disclosed. Methods of treating patients having or at risk of having AIDS or HIV infection, cancer, cardiac disorder, post-myocardial depression and post-traumatic stress disorder using optically pure R(-)fluoxetine in combination with one or more other biol. active compds. are further disclosed.
- AN 1999:763863 HCAPLUS <<LOGINID::20100601>>
- DN 132:6368
- ${\sf TI}$ Compositions and methods employing R(-)fluoxetine and other active ingredients
- IN Barberich, Timothy J.; Rubin, Paul D.; Handley, Dean A.
- PA Sepracor Inc., USA
- SO PCT Int. Appl., 41 pp. CODEN: PIXXD2
- DT Patent
- LA English

FAN.CNT 2

•	PA:	ATENT NO.)	DATE			APPLICATION NO.								
PI		9961014 9961014						19991202 20000720		WO 1999-US11725				19990527 <					
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	US	5 1998-177703				В2		1998	1023	<-	<								
	WO	O 1999-US11725 W 199				1999	.9990527 <												
	US 2000-664732 B3 20000919				0919	<-	_												
OSC.G		10 THERE ARE		10 CAPLUS RECORDS				S TH.	THAT CITE THIS RECORD					(10 CITINGS)					
RE.CNT		12	TH	ERE	ARE	12 CITED REFERENCES AVAILABLE FOR THIS RECORD													

- L15 ANSWER 39 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Combination of 5-HT3 receptor antagonist and serotonin reuptake inhibitor for treatment of depression

ALL CITATIONS AVAILABLE IN THE RE FORMAT

AB The present invention provides a method for treating a patient suffering from depression, comprising administering to said patient an effective amount of a first component which is a 5-HT3 receptor antagonist, in combination with an effective amount of a second component which is a serotonin reuptake inhibitor wherein improvement in sexual dysfunction and/or reduction in gastrointestinal side effects is recognized.

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Various formulations were prepared E.g., a tablet was prepared using
     zatosetron 10, fluoxetine HCl 10, microcryst. cellulose 275, fumed silica
     10, and stearic acid 5 mg, resp.
     1999:753081 HCAPLUS <<LOGINID::20100601>>
ΑN
DN
     131:346552
     Combination of 5-HT3 receptor antagonist and serotonin reuptake
TI
     inhibitor for treatment of depression
IN
     Michelson, David; Tollefson, Gary Dennis
PA
     Eli Lilly and Company, USA
SO
     PCT Int. Appl., 25 pp.
     CODEN: PIXXD2
DT
     Patent
     English
LA
FAN.CNT 1
     PATENT NO.
                          KIND DATE
                                            APPLICATION NO.
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     WO 9959593
                            A1 19991125 WO 1999-US10092
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RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                            A1 19991125 CA 1999-2332253
A 19991206 AU 1999-38912
A1 20010228 EP 1999-921795
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     JP 2002515435
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               THERE ARE 6 CAPLUS RECORDS THAT CITE THIS RECORD (8 CITINGS)
OSC.G 6
RE.CNT 7
               THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD
               ALL CITATIONS AVAILABLE IN THE RE FORMAT
L15 ANSWER 40 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
     Atypical antipsychotic agent-serotonin
ТΤ
     reuptake inhibitor combinations for therapy of refractory
     depression
AB
     Methods and compns. are provided for the treatment of depressive
     states refractory to treatment with traditional antidepressive therapies
     alone. These methods and compns. employ a compound having activity as an
     atypical antipsychotic (e.g. olanzapine) and a
     serotonin reuptake inhibitor (e.g. fluoxetine). This invention
     also provides methods of providing rapid onset treatments of major
     depression which employing a compound having activity as an
     atypical antipsychotic and a serotonin
     reuptake inhibitor.
     1999:752863 HCAPLUS <<LOGINID::20100601>>
AN
DN
     131:346550
ΤI
     Atypical antipsychotic agent-serotonin
     reuptake inhibitor combinations for therapy of refractory
     depression
     Tollefson, Gary Dennis
ΙN
PA
     Eli Lilly and Co., USA
SO
     Eur. Pat. Appl., 15 pp.
     CODEN: EPXXDW
DT
     Patent
LA
     English
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FAN.CNT 2
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    EP 958824 A2 19991124 EP 1999-303969 19990521 <-- 
EP 958824 A3 19991201
PΙ
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            THERE ARE 5 CAPLUS RECORDS THAT CITE THIS RECORD (5 CITINGS)
L15 ANSWER 41 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
     Methods for treating neuropsychiatric disorders
TI
     The invention provides methods for treating neuropsychiatric disorders
AΒ
     such as schizophrenia, Alzheimer's Disease, autism, depression,
     benign forgetfulness, childhood learning disorders, close head injury, and
     attention deficit disorder. The methods entail administering to a patient
     with a neuropsychiatric disorder a pharmaceutical composition containing (i) a
     therapeutically effective amount of D-alanine (or a modified form), provided
     that the composition is substantially free of D-cycloserine, and/or (ii)
     D-serine (or a modified form), and/or (iii) 105 to 500 mg of D-cycloserine
     (or a modified form), and/or (iv) N-methylglycine (or a modified form).
     Using double-blind conditions, patients were randomly assigned to receive
     placebo (fruit juice), D-serine 30, D-alanine 60-100, or N-methylglycine
     30 mg/kg/day once a day by mouth for 6 wk. Treatment with D-serine,
     D-alanine, or N-methylglycine improved the schizophrenic symptoms and
     cognitive deficit of the patients. Specifically, treatment with D-serine
     resulted in a 21% reduction of the neg. symptoms (on the SANS scale), and it
     resulted in a 17% reduction of the pos. symptoms. Treatment with D-alanine
     resulted in an 11% reduction of the neg. symptoms and a 12% reduction of the
pos.
     symptoms. Reatment with N-methylglycine resulted in a 20% reduction of the
     neg. symptoms and a 15% reduction of the pos. symptoms. These redns. in the
     neg. and pos. symptoms represented clin. significant improvement.
ΑN
    1999:672562 HCAPLUS <<LOGINID::20100601>>
DN
    131:281590
ΤI
    Methods for treating neuropsychiatric disorders
IN
     Tsai, Guochuan; Coyle, Joseph
PΑ
     The General Hospital Corporation, USA
SO
     PCT Int. Appl., 27 pp.
     CODEN: PIXXD2
DT
    Patent
LA
    English
FAN.CNT 1
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    EP 1844769
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    US 7704978
                       B2 20100427
    US 1998-81645P P 19980414 <--
US 1998-81654P P 19980414 <--
CA 1999-2328197 A3 19990414 <--
EP 1999-917453 A3 19990414 <--
PRAI US 1998-81645P
    US 1999-291296
                       A1 19990414 <--
    WO 1999-US8056
                       W
                             19990414 <--
    US 2001-834351
                       A1 20010413 <--
    US 2002-196686
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                        A1
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ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT
OSC.G 23
             THERE ARE 23 CAPLUS RECORDS THAT CITE THIS RECORD (24 CITINGS)
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L15 ANSWER 42 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN

RE.CNT 8

TI Controlled, double-blind investigation of the clozapine discontinuation symptoms with conversion to either olanzapine or placebo

ALL CITATIONS AVAILABLE IN THE RE FORMAT

THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD

AB The abrupt appearance of clozapine discontinuation symptoms represents a particularly unique situation that has not been characterized in a double-blind, placebo-controlled trial. A randomized, double-blind comparison of placebo (N = 53) and olanzapine 10 mg (N = 53) for 3 to 5 days following the abrupt discontinuation of clozapine (≤300 mg/day) was carried out. Subjects were assessed with the Pos. and Neg. Syndrome Scale (PANSS), the Clin. Global Impression Scale of Severity, the Montgomery-Asberg Depression Rating Scale (MADRS), and the Mini-Mental State Evaluation. Subsequently both groups received open-label olanzapine (10-25 mg/day) for an addnl. 9 wk. Statistically significantly more placebo-treated (24.5%) than olanzapine-treated (7.5%) patients experienced clozapine discontinuation symptoms (p = 0.017). Core symptoms included delusions, hallucinations, hostility, and paranoid

reaction and translated into a significantly higher worsening from baseline on the PANSS total, PANSS General Psychopathol. subscale, and MADRS among subjects randomly assigned to receive placebo. After open-label treatment with olanzapine for 9 wk, both groups were clin. stable, suggesting that the discontinuation symptoms were transient. However, subjects who had been randomly assigned to the 3- to 5-day placebo discontinuation segment achieved somewhat less global clin. improvement. Although a pharmacol. interpretation is speculative, evidence of a clozapine discontinuation syndrome was apparent. In most cases, the direct substitution of a pharmacol. similar agent (olanzapine) prevented the syndrome. Clozapine discontinuation or noncompliance should be considered in the differential assessment of an acutely emergent psychosis. The possibility that subjects who experience a clozapine discontinuation syndrome may take longer or are less likely to clin. restabilize warrants further investigation.

- AN 1999:657047 HCAPLUS <<LOGINID::20100601>>
- DN 131:266967
- TI Controlled, double-blind investigation of the clozapine discontinuation symptoms with conversion to either olanzapine or placebo
- AU Tollefson, Gary D.; Dellva, Mary Anne; Mattler, Carole A.; Kane, John M.; Wirshing, Donna A.; Kinon, Bruce J.; Ames, Donna; Cohn, Cal K.; Daniel, David G.; Clark, Scott C.; Horne, Robert L.; Kane, John M.; Levine, Robert; Miller, Marvin; Nemeroff, Charles B.; Reinstein, Michael R.; Smith, Thomas E.
- CS The Collaborative Crossover Study Group, Psychopharmacology Division, Lilly Research Laboratories, Lilly Corporate Center, Eli Lilly and Company, Indianapolis, IN, 46285, USA
- SO Journal of Clinical Psychopharmacology (1999), 19(5), 435-443 CODEN: JCPYDR; ISSN: 0271-0749
- PB Lippincott Williams & Wilkins
- DT Journal
- LA English
- OSC.G 9 THERE ARE 9 CAPLUS RECORDS THAT CITE THIS RECORD (9 CITINGS)
 RE.CNT 51 THERE ARE 51 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 43 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Is amoxapine an atypical antipsychotic?
 Positron-emission tomography investigation of its dopamine2 and serotonin2 occupancy
- AΒ All currently available atypical antipsychotics have, at clin. relevant doses: i) high serotonin (5-HT)2 occupancy; ii) greater 5-HT2 than dopamine (D)2 occupancy; and iii) a higher incidence of extrapyramidal side effects when their D2 occupancy exceeds 80%. A review of pharmacol. and behavioral data suggested that amoxapine should also conform to this profile; therefore, we undertook a positron-emission tomog. (PET) study of its 5-HT2 and D2 occupancy. Seven healthy volunteers received 50-250 mg/day of amoxapine for 5 days and then had [11C]-raclopride and [18F]-setoperone PET scans. 5-HT2 receptors showed near saturation at doses of 100 mg/day and above. The D2 receptor occupancies showed a dose-dependent increase, never exceeding 80%; at all doses 5-HT2 occupancy exceeded D2 occupancy. PET data show that amoxapine's profile is very similar to that of the established atypical antipsychotics. These data, together with amoxapine's in vitro pharmacol. profile, effectiveness in animal models, and efficacy in psychotic depression raise the possibility of amoxapine as an "atypical" antipsychotic agent in the treatment of schizophrenia.
- AN 1999:353093 HCAPLUS <<LOGINID::20100601>>
- DN 131:153673
- TI Is amoxapine an atypical antipsychotic?
 Positron-emission tomography investigation of its dopamine2 and serotonin2

occupancy

- AU Kapur, Shitij; Cho, Raymond; Jones, Corey; McKay, Gordon; Zipursky, Robert B.
- CS Center for Addictions and Mental Health, Clarke Institute of Psychiatry, Toronto, ON, M5T 1R8, Can.
- SO Biological Psychiatry (1999), 45(9), 1217-1220 CODEN: BIPCBF; ISSN: 0006-3223
- PB Elsevier Science Inc.
- DT Journal
- LA English
- OSC.G 12 THERE ARE 12 CAPLUS RECORDS THAT CITE THIS RECORD (12 CITINGS)
- RE.CNT 20 THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 44 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Risperidone augmentation of selective serotonin reuptake inhibitors in major depression
- AB Background: At low doses, risperidone acts as a 5-HT2 antagonist. Preclin. data suggest 5-HT2 antagonists may enhance the action of serotonin. This report examines the clin. use of risperidone to augment selective serotonin reuptake inhibitor (SSRI) antidepressants in patients who have not responded to SSRI therapy. Method: In 8 patients with major depressive disorder without psychotic features (DSM-IV) who had not responded to an SSRI, risperidone was added to the ongoing SSRI treatment. Hamilton Rating Scale for Depression scores were obtained before and after the addition of risperidone. Results: These 8 patients remitted within 1 wk of the addition of risperidone. Risperidone also appeared to have beneficial effects on sleep disturbance and sexual dysfunction. Conclusion: Risperidone may be a useful adjunct to SSRIs in the treatment of depression.
- AN 1999:293784 HCAPLUS <<LOGINID::20100601>>
- DN 130:332801
- TI Risperidone augmentation of selective serotonin reuptake inhibitors in major depression
- AU Ostroff, Robert B.; Nelson, J. Craig
- CS Spectrum Psychiatric Group, P.C., Hamden, Conn., Hamden, CT, 06518, USA
- SO Journal of Clinical Psychiatry (1999), 60(4), 256-259 CODEN: JCLPDE; ISSN: 0160-6689
- PB Physicians Postgraduate Press, Inc.
- DT Journal
- LA English
- OSC.G 102 THERE ARE 102 CAPLUS RECORDS THAT CITE THIS RECORD (102 CITINGS)
- RE.CNT 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 45 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Olanzapine response in psychotic depression
- AB Psychotic depression is more common than is generally realized, occurring in an estimated 16% to 54% of depressed patients. In controlled studies of patients with schizophrenia, the atypical antipsychotic olanzapine has been shown to be superior in efficacy to haloperidol at doses of 10 mg/day. Since olanzapine may have antidepressant effects in addition to its antipsychotic properties, the purpose of this study was to assess the safety and efficacy of olanzapine in the treatment of psychotic depression. Hospitalized patients with the discharge diagnosis of DSM-IV psychotic depression (major depression with psychotic features or bipolar I disorder, depressed phase...with psychotic features) who had been treated with olanzapine during the first 9 mo of its availability in the United States were identified. An age- and sex-matched sample of

hospitalized patients with psychotic depression treated with other antipsychotics during the same time period was also identified. The medical records were expunged of all refs. to medication treatment and then reviewed and scored in a blind fashion for indications, doses, response, and side effects. Fifteen psychotic depression patients (10 women, 5 men), aged 36.9 ± 10.1 yr, who were treated with olanzapine were retrospectively compared with 15 psychotic depression patients (10 women, 5 men), aged $35.0 \pm 8.2 \text{ yr}$, treated with other antipsychotics. Ten (67%) of 15 patients taking olanzapine were much or very much improved upon discharge compared with only 4 (27%) of 15 patients taking other antipsychotics (Fisher exact test, p = .037). Olanzapine was well tolerated: no patient discontinued the medication because of side effects. Twelve (80%) of 15 patients in each group were taking antidepressants in addition to the antipsychotic. Of the 3 patients taking olanzapine but not taking an antidepressant , 2 were much or very much improved (1 patient taking olanzapine alone, 1 taking olanzapine plus valproate sodium). Olanzapine appears to be effective and safe for patients with psychotic depression. Further prospective studies are warranted to ascertain whether olanzapine's unique pharmacol. profile may make it particularly useful for the treatment of psychotic depression either alone or in combination with antidepressants.

- AN 1999:180184 HCAPLUS <<LOGINID::20100601>>
- DN 130:262002
- TI Olanzapine response in psychotic depression
- AU Rothschild, Anthony J.; Bates, Kimberly S.; Boehringer, Kelly L.; Syed,
- CS Department of Psychiatry, University of Massachusetts Medical School, Worcester, MA, USA
- SO Journal of Clinical Psychiatry (1999), 60(2), 116-118 CODEN: JCLPDE; ISSN: 0160-6689
- PB Physicians Postgraduate Press, Inc.
- DT Journal
- LA English
- OSC.G 37 THERE ARE 37 CAPLUS RECORDS THAT CITE THIS RECORD (37 CITINGS)
- RE.CNT 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 46 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Switching from clozapine to olanzapine in treatment-refractory schizophrenia: safety, clinical efficacy, and predictors of response
- AB In our experience, many of our schizophrenic patients treated with clozapine request the newer atypical antipsychotic agents in order to eliminate the weekly blood monitoring. However, there are few guidelines available to clinicians interested in switching patients successfully treated with clozapine to olanzapine. The goal of this study was to collect preliminary data on the safety, clin. effectiveness, and predictors of response of switching clozapine patients to olanzapine. In an open trial, 19 patients receiving clozapine were switched to olanzapine. Eight (42%) of 19 patients were considered responders. Seven patients decompensated seriously enough to require hospitalization. All 7 of these patients were restabilized on clozapine treatment in the hospital, and olanzapine was discontinued. In an addnl. 4 patients, clin. status worsened, and clozapine doses were titrated upwards and olanzapine was slowly discontinued. Overall, mean total Brief Psychiatric Rating Scale (BPRS) scores increased significantly from baseline to final assessment (p = .02). Responders had been treated for a significantly shorter period of time with clozapine prior to the switch compared to nonresponders (p = .04) and were receiving a lower dose of clozapine (p = .05). The final olanzapine dose did not differ between responders and nonresponders. All responders have remained on olanzapine

treatment and are stable. In this open trial, the crossover from clozapine to olanzapine was generally well tolerated and resulted in a successful transition for 8 of the 19 patients. However, mean scores on the total BPRS and neg. symptom and depressive symptom subscales significantly increased. Caution must be taken in determining which patients may benefit from the switch to olanzapine because of the risk of decompensation and hospitalization. Because this was an open trial, these findings require replication in a controlled trial.

- AN 1999:6903 HCAPLUS <<LOGINID::20100601>>
- DN 130:218117
- TI Switching from clozapine to olanzapine in treatment-refractory schizophrenia: safety, clinical efficacy, and predictors of response
- AU Henderson, David C.; Nasrallah, Rima A.; Goff, Donald C.
- CS Erich Lindemann Mental Health Center and the Psychiatry Service, Massachusetts General Hospital, Boston, USA
- SO Journal of Clinical Psychiatry (1998), 59(11), 585-588 CODEN: JCLPDE; ISSN: 0160-6689
- PB Physicians Postgraduate Press, Inc.
- DT Journal
- LA English
- OSC.G 17 THERE ARE 17 CAPLUS RECORDS THAT CITE THIS RECORD (17 CITINGS)
- RE.CNT 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 47 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI A double-blind, placebo-controlled study of risperidone in adults with autistic disorder and other pervasive developmental disorders
- AΒ Neurobiol. research has implicated the dopamine and serotonin systems in the pathogenesis of autism. Open-label reports suggest that the serotonin2A-dopamine D2 antagonist risperidone may be safe and effective in reducing the interfering symptoms of patients with autism. Thirty-one adults (age [mean + SD], 28.1 ± 7.3 yr) with autistic disorder (n = 17) or pervasive developmental disorder not otherwise specified (n = 14) participated in a 12-wk double-blind, placebo-controlled trial of risperidone. Patients treated with placebo subsequently received a 12-wk open-label trial of risperidone. For persons completing the study, 8 (57%) of 14 patients treated with risperidone were categorized as responders (daily dose [mean ± SD], 2.9 \pm 1.4 mg) compared with none of 16 in the placebo group (P<.002). Risperidone was superior to placebo in reducing repetitive behavior (P<.001), aggression (P<.001), anxiety or nervousness (P<.02), depression (P<.03), irritability (P<.01), and the overall behavioral symptoms of autism (P<.02). Objective, measurable change in social behavior and language did not occur. Nine (60%) of 15 patients who received treatment with open-label risperidone following the double-blind placebo phase responded. Other than mild, transient sedation, risperidone was well tolerated, with no evidence of extrapyramidal effects, cardiac events, or seizures. Risperidone is more effective than placebo in the short-term treatment of symptoms of autism in adults.
- AN 1998:482360 HCAPLUS <<LOGINID::20100601>>
- DN 129:254836
- OREF 129:51743a,51746a
- TI A double-blind, placebo-controlled study of risperidone in adults with autistic disorder and other pervasive developmental disorders
- AU Mcdougle, Christopher J.; Holmes, Janice P.; Carlson, Derek C.; Pelton, Gregory H.; Cohen, Donald J.; Price, Lawrence H.
- CS Department of Psychiatry, Section of Child and Adolescent Psychiatry, Indiana University School of Medicine, Indianapolis, USA
- SO Archives of General Psychiatry (1998), 55(7), 633-641 CODEN: ARGPAQ; ISSN: 0003-990X
- PB American Medical Association

DT Journal LA English

OSC.G 56 THERE ARE 56 CAPLUS RECORDS THAT CITE THIS RECORD (56 CITINGS)

RE.CNT 38 THERE ARE 38 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L15 ANSWER 48 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN

- TI A double-blind, controlled comparison of the novel antipsychotic olanzapine versus haloperidol or placebo on anxious and depressive symptoms accompanying schizophrenia
- Depressive symptoms are a common feature of schizophrenia and AB may represent a core part of the illness. Where present, it has been associated with greater overall morbidity and mortality. Monotherapy with conventional dopamine antagonists may either worsen or bestow a limited therapeutic benefit. Accordingly the use of adjunctive thymoleptics has been explored. In contrast, olanzapine (OLZ), an atypical antipsychotic agent, offers a distinctive and pleiotropic pharmacol. suggestive of a broader efficacy profile than conventional neuroleptic agents. In a 6-wk placebo- and haloperidol (HAL)-controlled trial with 335 randomized subjects with chronic schizophrenia in an acute exacerbation, three fixed dose ranges of OLZ $(5, 10, or 15 \pm 2.5 mg)$ were evaluated vs. HAL (10-20 mg) or placebo. Baseline to endpoint change in the Brief Psychiatric Rating Scale including the anxietydepression cluster (items 1, 2, 5, 9) was analyzed. Two dose ranges of OLZ (10 \pm 2.5, 15 \pm 2.5) were superior to placebo (p < .05) in improving mood status, whereas HAL was not. Contributions from a more selective meso-limbic dopaminergic profile, D1 or D4 activity, the release of dopamine/norepinephrine in the prefrontal cortex, and/or serotonin 5-HT2A,C antagonism may explain the differential benefit seen with OLZ in the treatment of comorbid anxious and depressive symptoms in schizophrenia.
- AN 1998:373967 HCAPLUS <<LOGINID::20100601>>

DN 129:117695

OREF 129:23985a,23988a

- TI A double-blind, controlled comparison of the novel antipsychotic olanzapine versus haloperidol or placebo on anxious and depressive symptoms accompanying schizophrenia
- AU Tollefson, Gary D.; Sanger, Todd M.; Beasley, Charles M.; Tran, Pierre V.
- CS Lilly Research Laboratories, Eli Lilly and Company, Indianapolis, IN, 46285, USA
- SO Biological Psychiatry (1998), 43(11), 803-810 CODEN: BIPCBF; ISSN: 0006-3223
- PB Elsevier Science Inc.
- DT Journal
- LA English
- OSC.G 41 THERE ARE 41 CAPLUS RECORDS THAT CITE THIS RECORD (41 CITINGS)
- RE.CNT 53 THERE ARE 53 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 49 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Thyroid parameters during therapy with zotepine in delusional depression: preliminary results
- AB There are only few data on the effects of atypical neuroleptics on thyroid function. In an open pilot study of 12 inpatients with delusional depression, thyroid hormone levels and TRH-TSH test were determined during neuroleptic treatment with zotepine. No significant changes in triiodothyronine (T3), thyroxine (T4) and delta-TSH levels were found in this observation period (28 days).
- AN 1998:266943 HCAPLUS <<LOGINID::20100601>>
- DN 129:23598
- OREF 129:4915a,4918a

- TI Thyroid parameters during therapy with zotepine in delusional depression: preliminary results
- AU Konig, Frank; Hauger, Barbara; Barg, Thomas; Wolfersdorf, Manfred
- CS Depression Unit, Weissenau Psychiatric Center, Department of Psychiatry I, University of Ulm, Ravensburg, D-88214, Germany
- SO Neuropsychobiology (1998), 37(2), 88-90 CODEN: NPBYAL; ISSN: 0302-282X
- PB S. Karger AG
- DT Journal
- LA English
- OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)
- RE.CNT 23 THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 50 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Blood biogenic amines during clozapine treatment of early-onset schizophrenia
- AΒ The aims of this investigation were to evaluate long-term and short-term effects of clozapine-treatment on plasma biogenic amines and psychopathol. measures in adolescents with schizophrenia (DSM-III-R criteria). The long-term study was conducted in a study sample of 40 young patients (age 14-22 yr) following a mean of 3.4 yr of neuroleptic treatment. During the study, 20 patients received clozapine, and the other 20 patients were treated with standard neuroleptic medications. At the beginning of the open clin. trials, the patients had already been receiving clozapine treatment for 24 + 15 mo. Assessment of the biochem. and psychopathol. measures was performed on six occasions at consecutive 6-wk intervals during maintenance treatment with clozapine or conventional neuroleptics. Blood levels of serotonin, 3-methoxy-4-hydroxy-phenylglycol (MHPG), norepinephrine, and epinephrine were significantly higher in clozapine-treated patients than in conventionally treated patients. During long-term treatment, higher serotonin levels were associated with significantly fewer neg. symptoms of schizophrenia, whereas higher MHPG levels were correlated with less depression. The short-term effects of clozapine were assessed in a second and independent study sample. After failing on conventional neuroleptics in clin. trials lasting a mean of 1.6 yr, 15 inpatients (aged 11-20 yr) received clozapine. Weekly ratings of psychopathol. symptoms using standard rating scales were performed in parallel to blood samplings for measurements of biogenic amines and serum levels of clozapine. These measures were obtained for 6 wk during conventional neuroleptic treatment and for 6 wk during the open-label clozapine trial. Serum levels of serotonin and plasma norepinephrine levels were significantly higher during treatment with clozapine than during pretreatment with typical neuroleptics. A comparison of plasma epinephrine levels in responders (n = 7) and nonresponders (n = 8) to clozapine revealed that response to clozapine can be predicted by epinephrine levels prior to initiation of treatment with clozapine (responders ranging from 32.2 to 90.3 pg/mL; nonresponders ranging from 92.5 to 473.5 pg/mL). Addnl., subjects who responded to clozapine showed increased mean plasma concns. of MHPG and epinephrine during treatment with this drug in comparison to the levels measured during pretreatment with typical neuroleptic medication. Nonresponders to clozapine failed to show this increase. Finally, in responders to clozapine a neg. linear relationship between neg. symptoms of schizophrenia and the concns. of plasma norepinephrine and serum plasma epinephrine levels prior to initiation of clozapine therapy predict response to this atypical neuroleptic. Our findings derived from short-term and maintenance treatment with clozapine suggest involvement of norepinephrine, epinephrine and serotonin in the therapeutic actions of the atypical neuroleptic clozapine.

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AN 1998:144787 HCAPLUS <<LOGINID::20100601>>
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DN 128:239382

OREF 128:47237a,47240a

TI Blood biogenic amines during clozapine treatment of early-onset schizophrenia

AU Schulz, E.; Fleischhaker, C.; Clement, H.-W.; Remschmidt, H.

CS Departments of 'Child and Adolescent Psychiatry, Philipps-University, Marburg, Germany

SO Journal of Neural Transmission (1997), 104(10), 1077-1089 CODEN: JNTRF3; ISSN: 0300-9564

PB Springer-Verlag Wien

DT Journal

LA English

OSC.G 10 THERE ARE 10 CAPLUS RECORDS THAT CITE THIS RECORD (10 CITINGS)

RE.CNT 52 THERE ARE 52 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L15 ANSWER 51 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN

TI The effects of risperidone on the five dimensions of schizophrenia derived by factor analysis: combined results of the North American trials

AΒ In 2 double-blind trials conducted in North America, 513 patients with chronic schizophrenia received risperidone, haloperidol, or placebo. the present study, combined data from the 2 trials were analyzed. Patients were randomly assigned to receive placebo, risperidone (2, 6, 10, and 16 mg/day), or haloperidol (20 mg/day) for 8 wk. Factor anal. of scores on the Pos. and Neq. Syndrome Scale (PANSS) produced 5 dimensions (neg. symptoms, pos. symptoms, disorganized thought, uncontrolled hostility/excitement, and anxiety/depression), similar to the 5 dimensions of previous factor-analytic studies of PANSS data. Mean changes (symptom redns.) in PANSS factor scores from basal values to treatment weeks 6 and 8 were greater in patients receiving 6-16 mg risperidone/day than in patients receiving placebo or haloperidol. advantages of risperidone were greatest for neg. symptoms, uncontrolled hostility/excitement, and anxiety/depression. Even at the lowest dose, 2 mg/day, risperidone was superior to haloperidol in reducing neg. symptoms. The differences in outcomes between the effects of risperidone and haloperidol on PANSS scores were not related to extrapyramidal symptoms. Risperidone produced greater improvements than haloperidol on all 5 dimensions. The large between-group differences in effect on neg. symptoms, hostility/excitement, and anxiety/ depression suggest that risperidone and other serotonin /dopamine antagonists have qual. different effects from those of conventional antipsychotic agents.

AN 1998:60811 HCAPLUS <<LOGINID::20100601>>

DN 128:200905

OREF 128:39583a,39586a

TI The effects of risperidone on the five dimensions of schizophrenia derived by factor analysis: combined results of the North American trials

AU Marder, Stephen R.; Davis, John M.; Chouinard, Guy

CS West Los Angeles Veterans Affairs Medical Center, Los Angeles, CA, 90037, USA

SO Journal of Clinical Psychiatry (1997), 58(12), 538-546 CODEN: JCLPDE; ISSN: 0160-6689

PB Physicians Postgraduate Press

DT Journal

LA English

OSC.G 104 THERE ARE 104 CAPLUS RECORDS THAT CITE THIS RECORD (104 CITINGS)

RE.CNT 33 THERE ARE 33 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- TI Olanzapine-pharmacology and clinical evaluation of a new atypical antipsychotic
- AB A review with 47 refs. Olanzapine is one of a number of newer " atypical" antipsychotic agents which are emerging from attempts to find better tolerated and more effective drugs in the treatment of schizophrenia. It has structural and pharmacol. similarities to the atypical antipsychotic clozapine, the first agent to demonstrate significant therapeutic advantages over standard antipsychotic agents. Preclin. studies found olanzapine to have a broad range of receptor affinities including actions at dopamine and serotonin receptors and were also predictive of its atypical antipsychotic potential. Double-blind controlled trials involving more than 2900 patients have shown it to be an effective antipsychotic which induces low levels of extrapyramidal adverse effects, justifying its atypical status. Addnl. benefits suggested include efficacy in reducing neg. symptoms and a favorable effect on comorbid depressive symptoms. It has been found to be well tolerated with a relatively low tendency to cause sustained elevations in serum prolactin levels. No major adverse events have been reported. While olanzapine appears to offer advantages compared with standard antipsychotics, use in clin. practise and further trials are required to clarify its full therapeutic potential.

AN 1997:738168 HCAPLUS <<LOGINID::20100601>>

DN 128:29960

OREF 128:5725a,5728a

- TI Olanzapine-pharmacology and clinical evaluation of a new atypical antipsychotic
- AU Weaver, Mark G.
- CS Department of Psychological Medicine, St Bartholomew's Hospital, London, EC1A 7BE, UK
- SO Journal of Serotonin Research (1997), 4(2), 145-157 CODEN: JSRRER; ISSN: 1350-7702
- PB Euroscience Press
- DT Journal; General Review
- LA English
- OSC.G 4 THERE ARE 4 CAPLUS RECORDS THAT CITE THIS RECORD (4 CITINGS)
- RE.CNT 47 THERE ARE 47 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L15 ANSWER 53 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Ziprasidone
- AB A review with 24 refs. Ziprasidone is a novel antipsychotic drug. It has high affinity for serotonin 5-HT2 and dopamine D2 receptors in vitro, with an 11-fold higher affinity for 5-HT2 than for D2 receptors, suggestive of a low potential for inducing motor disturbance [including extrapyramidal symptoms (EPS)]. The effects of ziprasidone in receptor binding studies reflected its in vitro pharmacol., with more potent effects against 5-HT2 receptor-than against D2 receptor-mediated behavior. Because ziprasidone inhibits serotonin (5-hydroxytryptamine; 5-HT) and noradrenaline (norepinephrine) reuptake, it may have anxiolytic and antidepressant effects. Data from phase II and III clin. trials have shown ziprasidone to be effective in reducing the pos. and neg. symptoms of, and depression associated with, schizophrenia, and in reducing anxiety in patients about to undergo dental surgery. Ziprasidone was generally well tolerated in phase II and III clin. trials, with somnolence and nausea being the most frequently reported adverse events in placebo-controlled studies. Motor disturbances, including EPS, were infrequently observed

AN 1997:593623 HCAPLUS <<LOGINID::20100601>>

DN 127:242699

OREF 127:47191a,47194a

```
Ziprasidone
ΤI
     Davis, Rick; Markham, Anthony
ΑU
CS
     Adis International Limited, Auckland, N. Z.
SO
     CNS Drugs (1997), 8(2), 153-159
     CODEN: CNDREF; ISSN: 1172-7047
PΒ
     Adis
DT
     Journal; General Review
LA
     English
              THERE ARE 15 CAPLUS RECORDS THAT CITE THIS RECORD (15 CITINGS)
OSC.G
      15
L15 ANSWER 54 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
     Dosing the antipsychotic medication olanzapine
AΒ
     A review with 5 refs. Olanzapine is a new antipsychotic agent with
     serotonin/dopamine antagonism action. Efficacy in treating
     overall psychopathol. in acute schizophrenia as measured by the BPRS0-6
     total score was demonstrated at 10 mg/day vs. placebo; at doses in a 5-20
     mg/day range, olanzapine was comparable or superior to haloperidol.
     Superior efficacy for neg. and depressive symptoms was shown in
     comparison to haloperidol. Olanzapine has a favorable acute and tardive
     extrapyramidal symptom profile relative to haloperidol and caused
     substantially less elevation of serum prolactin. Dose-related weight gain
     and asymptomatic mild transaminase elevations occurred in
     olanzapine-treated patients.
     1997:567795 HCAPLUS <<LOGINID::20100601>>
ΑN
DN
     127:214477
OREF 127:41537a
    Dosing the antipsychotic medication olanzapine
TΙ
ΑU
    Nemeroff, Charles B.
CS
     Department of Psychiatry and Behavioral Sciences, Emory University School
     of Medicine, Atlanta, GA, 30322, USA
SO
     Journal of Clinical Psychiatry (1997), 58 (Suppl. 10), 45-49
    CODEN: JCLPDE; ISSN: 0160-6689
PΒ
    Physicians Postgraduate Press
DT
     Journal; General Review
     English
LA
OSC.G
       9
              THERE ARE 9 CAPLUS RECORDS THAT CITE THIS RECORD (9 CITINGS)
L15 ANSWER 55 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
    Method for treating depression
ΤI
     The invention provides a method for treating depressive signs
AΒ
     and symptoms comprising administering an effective amount of
     2-methyl-4-(4-methyl-1-piperazinyl)-10H-thieno[2,3-b][1,5]benzodiazepine
     to a patient in need thereof.
ΑN
    1997:503273 HCAPLUS <<LOGINID::20100601>>
    127:126642
DN
OREF 127:24313a,24316a
    Method for treating depression
ΤI
     Tollefson, Gary D.
ΙN
     Eli Lilly and Company, USA; Tollefson, Gary D.
PΑ
SO
     PCT Int. Appl., 11 pp.
     CODEN: PIXXD2
DT
     Patent
LA
    English
FAN.CNT 1
                       KIND
                               DATE
                                          APPLICATION NO.
     PATENT NO.
                        A1 19970703
                                         WO 1996-US19574 19961204 <--
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            LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT,
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                              19981007
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                              19990120 CN 1996-199221
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                       А3
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    NZ 325036
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    US 5958921
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PRAI US 1995-9173P
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ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT
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OSC.G

RE.CNT 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L15 ANSWER 56 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- ΤI Risperidone dose-dependently increases extracellular concentrations of serotonin in the rat frontal cortex: role of $\alpha 2$ -adrenoceptor antagonism
- We have previously shown that risperidone, an antipsychotic drug with high AB affinity for 5-hydroxytryptamine (5-HT)2A and dopamine (DA)2 receptors, as well as for $\alpha 1$ -and $\alpha 2$ -adrenoceptors, enhances 5-HT metabolism selectivity in the rat frontal cortex (FC). To further study the influence of risperidone on central 5-HT systems, we compared its effects on dialyzate 5-HT in the FC, as assessed by microdialysis, with those obtained with other antipsychotic drugs, i.e., clozapine, haloperidol, and amperozide, as well as with the selective $\alpha 2-$ or 5-HT2A receptor antagonists idazoxan or MDL 100,907, resp. The underlying mechanism for risperidone's effect on 5-HT output in the FC was also investigated using single-cell recording in the dorsal raphe nucleus (DRN). Administration of risperidone (0.2, 0.6, and 2.0 mg/kg, SC) dose-dependently increased 5-HT levels in the FC. This stimulatory action was mimicked by amperozide (10 mg/kg, SC) and, to some extent, by idazoxan (0.25 mg/kg, SC). In contrast, clozapine (10 mg/kg, SC), haloperidol (2.0 mg/kg, SC), and MDL 100,907 (1.0 mg/kg, SC) exerted only minor effects on 5-HT output in brain. Local administration of risperidone or idazoxan (1.0-1000 $\mu\text{mol/L})$ in the FC dose-dependently increased dialyzate levels of 5-HT in this region. On the other hand, risperidone (25-800 $\mu g/kg$, IV) dose-dependently decreased the firing rate of 5-HT cells in the DRN, an effect that was largely antagonized by pretreatment with the selective 5-HT1A receptor antagonist WAY 100,635 (5.0 μ g/kg, IV). These results indicate that the risperidone-increased 5-HT output in the FC may be related to its α 2-adrenoceptor antagonistic action, a property shared with both amperozide and idazoxan, and that this action probably is executed at the nerve terminal level. The inhibition of 5-HT cell firing by risperidone is probably secondary to increased 5-HT availability, e.g., in the DRN, since it could be antagonized by a 5-HT1A receptor antagonist. The enhanced 5-HT output in the FC by risperidone may be of particular relevance for the treatment of schizophrenia when associated with depression and in schizoaffective disorder.
- ΑN 1997:408932 HCAPLUS <<LOGINID::20100601>>
- DN 127:90420
- OREF 127:17233a,17236a
- Risperidone dose-dependently increases extracellular concentrations of serotonin in the rat frontal cortex: role of $\alpha 2$ -adrenoceptor

antagonism

AU Hertel, Peter; Nomikos, George G.; Schilstroem, Bjoern; Arborelius, Lotta; Svensson, Torgny H.

CS Department of Physiology and Pharmacology, Division of Pharmacology, Karolinska Institutet, Stockholm, S-171 77, Swed.

SO Neuropsychopharmacology (1997), 17(1), 44-55 CODEN: NEROEW; ISSN: 0893-133X

PB Elsevier

DT Journal

LA English

OSC.G 44 THERE ARE 44 CAPLUS RECORDS THAT CITE THIS RECORD (44 CITINGS)

RE.CNT 62 THERE ARE 62 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L15 ANSWER 57 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN

 ${\tt TI}$ Preparation of (piperazinyl)dibenzoxazepines as 5-HT2 receptor ligands ${\tt GI}$

AB The piperazine derivs, I (A, B = ring-forming group; X1 = 0, S, etc.; X2 = imino, methine, carbonyl, etc.; R1 = alkyl, etc.; R2, R3, R4 = H, alkyl) were disclosed as 5-HT2 receptor-selective compds. The compds. I are analogs of clozapine. The use of I in the serotonin 5-HT2 receptor identification and use in drug screening programs and as pharmaceuticals to treat indications in which the 5-HT2 receptor is implicated, such as hypertension, thrombosis, migraine, vasospasm, ischemia, depression, anxiety, schizophrenia, sleep disorders and appetite disorders were also described.

AN 1996:473238 HCAPLUS <<LOGINID::20100601>>

DN 125:142796

OREF 125:26741a

TI Preparation of (piperazinyl)dibenzoxazepines as 5-HT2 receptor ligands

IN Tehim, Ashok; Fu, Jian-Min; Rakhit, Sumanas

PA Allelix Biopharmaceuticals Inc., Can.

SO PCT Int. Appl., 27 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

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MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI,
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             IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR,
             NE, SN, TD, TG
     US 5602124
                                19970211
                                           US 1994-354765
                                                                   19941212 <--
                         Α
     CA 2207613
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                                           CA 1995-2207613
                                                                   19951208 <--
     AU 9539348
                         Α
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PRAI US 1994-354765
                                19941212
                         Α
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     WO 1995-IB1111
                                19951208 <--
                         W
ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT
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              THERE ARE 13 CAPLUS RECORDS THAT CITE THIS RECORD (18 CITINGS)
OSC.G
      13
RE.CNT 3
              THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
    ANSWER 58 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
L15
ΤI
     Risperidone therapy in treatment refractory acute bipolar and
     schizoaffective mania
```

- AΒ This pilot study evaluated the efficacy of risperidone therapy in patients with bipolar I or schizoaffective mania who were treatment resistant or treatment intolerant. Patient psychopathol. and involuntary movements were evaluated with a variety of scales, and risperidone was administered on an open-label basis. Five of six patients (all bipolar) discontinued risperidone therapy because of adverse drug effects (2 patients), lack of significant drug response and subjective clin. worsening (1 patient), or worsening of manic symptoms (2 patients). One patient with schizoaffective illness improved. Risperidone used without the addition of a mood stabilizer was ineffective in treating pure manic psychosis. In some vulnerable bipolar patients, risperidone monotherapy may have antidepressant activity that could exacerbate mania. If risperidone proves to have antidepressant activity, it may become an important agent in the therapy of patients with depressive symptoms and psychosis.
- AN 1996:426696 HCAPLUS <<LOGINID::20100601>>
- DN 125:76241
- OREF 125:14275a,14278a
- TI Risperidone therapy in treatment refractory acute bipolar and schizoaffective mania
- AU Sajatovic, Martha; DiGiovanni, Sue Kim; Bastani, Bijan; Hattab, Helen; Ramirez, Luis F.
- CS Medical Center, Cleveland Veterans Administration, Cleveland, OH, 44141, USA
- SO Psychopharmacology Bulletin (1996), 32(1), 55-61 CODEN: PSYBB9; ISSN: 0048-5764
- PB U.S. Dep. of Health and Human Services
- DT Journal
- LA English
- OSC.G 29 THERE ARE 29 CAPLUS RECORDS THAT CITE THIS RECORD (29 CITINGS)
- L15 ANSWER 59 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Risperidone: regional effects in vivo on release and metabolism of dopamine and serotonin in the rat brain
- AB The antipsychotic drug risperidone shows high affinity for both central serotonin (5-HT)2A and dopamine (DA)-D2 receptors in vivo. By employing microdialysis in freely moving rats, the effects of acute risperidone administration on regional brain DA and 5-HT release and metabolism were compared with the corresponding effects of the atypical antipsychotic drug clozapine as well as amperozide, the selective DA-D2 receptor antagonist raclopride and the

selective 5-HT2A/5-HT2C receptor antagonist ritanserin. Risperidone (0.2 or 2.0 mg/kg, SC) was found to increase DA release and metabolism to about the same extent in three major projection areas of the mesotelencephalic dopaminergic system, i.e. the nucleus accumbens (NAC), the medial prefrontal cortex (MPC) and the lateral striatum (STR). In contrast, clozapine and amperozide (both 10.0 mg/kg, SC), as well as raclopride (2.0 mg/kg, SC), were all found differentially to affect DA release and metabolism in three projections areas. Specifically, clozapine and amperozide enhanced DA release in the MPC to a greater extent than in the NAC or the STR, whereas raclopride instead preferentially increased DA release in the NAC and the STR but not the MPC. Ritanserin (3.0 mg/kg, SC) did not exert any major effects on DA metabolism in the three areas studied. In contrast to the regionally rather homogeneous activation of brain DA systems caused by risperidone, the drug was found to enhance brain 5-HT metabolism preferentially in the MPC, as indicated by the elevated extracellular concentration of 5-hydroxyindoleacetic acid (5-HIAA) in this region. A similar elevation of the 5-HIAA level in the MPC was observed after amperozide and, to some extent, after clozapine and ritanserin administration. The risperidone-induced (2.0 mg/kg, SC) elevation of 5-HIAA concns. in the frontal cortex was paralleled by an increased 5-HT release in brain area. Consequently, the authors' findings demonstrated pharmacol. profile of risperidone, as reflected brain DA metabolism, in between that of clozapine and the Da-D2 antagonists. The preferential activation of 5-HT release and metabolism in frontal cortical areas might be of particular relevance for the ameliorating effect of risperidone on neg. symptoms in schizophrenia, especially when associated with depression.

AN 1996:355768 HCAPLUS <<LOGINID::20100601>>

DN 125:49065

OREF 125:9185a,9188a

- TI Risperidone: regional effects in vivo on release and metabolism of dopamine and serotonin in the rat brain
- AU Hertel, Peter; Nomikos, Geroge G.; Iurlo, Marina; Svensson, Torgny H.
- CS Dep. Physiol. Pharmacol., Karolinska Inst., Stockholm, S-171 77, Swed.
- SO Psychopharmacology (Berlin) (1996), 124(1/2), 74-86 CODEN: PSCHDL; ISSN: 0033-3158
- PB Springer
- DT Journal
- LA English
- OSC.G 73 THERE ARE 73 CAPLUS RECORDS THAT CITE THIS RECORD (74 CITINGS)
- L15 ANSWER 60 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI $\alpha 2$ -Adrenoreceptor antagonism may contribute to the atypical properties of risperidone: Experimental support for the Nutt case
- AB The therapeutic efficacy and reduction in side effects claimed for new antischizophrenic drugs such as clozapine and risperidone have been ascribed to their heightened affinity for serotonin 5-HT2 receptors rather than D-2 receptors. D. J. Nutt (1994) questioned the proposed role of 5-HT2 receptors, and focused instead on the possible contribution of $\alpha 2$ -adrenoreceptor antagonism. The authors have confirmed that at least one atypical property of risperidone (a rapid decrement in its ability to depress self-stimulation) can be partly prevented by an $\alpha 2$ -adrenoreceptor agonist (clonidine) but not by a 5-HT2 receptor agonist (DOI). This result supports the suggested role of $\alpha 2$ -adrenoreceptor antagonism in counteracting extrapyramidal effects during treatment with risperidone.
- AN 1995:940156 HCAPLUS <<LOGINID::20100601>>
- DN 124:45460
- OREF 124:8351a,8354a
- TI $\alpha2\text{--}Adrenore ceptor antagonism may contribute to the atypical properties of risperidone: Experimental support for the Nutt case$
- AU Herberg, L. J.; Montgomery, A. M. J.; Grottick, A. J.

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CS Institute Neurology, London, WC1N 3BG, UK
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- SO Journal of Psychopharmacology (Oxford) (1995), 9(3), 281-3 CODEN: JOPSEQ; ISSN: 0269-8811
- PB Oxford University Press
- DT Journal
- LA English
- OSC.G 7 THERE ARE 7 CAPLUS RECORDS THAT CITE THIS RECORD (7 CITINGS)
- L15 ANSWER 61 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- ${\tt TI}$ Clozapine acts as a 5-HT2 antagonist by attenuating DOI-induced inhibition of male rat sexual behavior
- AB Evidence has been reported that clozapine may derive part of its therapeutic effects in treatment of resistant schizophrenic patients by interacting with the serotonin system. Among the few behavioral models available to test the hypothesis of an interaction of clozapine with 5-HT2 receptors, male rat sexual behavior is particularly useful, since in this behavior 5-HT1A and 5-HT2 receptors have opposite functions. Stimulation of 5-HT1A receptors facilitates ejaculatory behavior and stimulation of 5-ZHT2 receptors inhibit ejaculation. In the present study, male rat sexual behavior was depressed by treatment with DOI (1.0 mg/kg), a selective 5-HT2 receptor agonist. The depressive effect of DOI was attenuated by the administration of clozapine (0.1-1.0 mg/kg) in doses that by themselves did not significantly affect sexual behavior. It was concluded that clozapine in the male rat sexual behavior model may be interpreted as serving as a 5-HT2 antagonist.
- AN 1995:666209 HCAPLUS <<LOGINID::20100601>>
- DN 123:74764
- OREF 123:13031a,13034a
- ${\tt TI}$ Clozapine acts as a 5-HT2 antagonist by attenuating DOI-induced inhibition of male rat sexual behavior
- AU Klint, T.; Larsson, K.
- CS Dep. of Psychology, Univ. of Goeteborg, Goeteborg, S-41314, Swed.
- SO Psychopharmacology (Berlin) (1995), 119(3), 291-4 CODEN: PSCHDL; ISSN: 0033-3158
- PB Springer
- DT Journal
- LA English
- OSC.G 19 THERE ARE 19 CAPLUS RECORDS THAT CITE THIS RECORD (19 CITINGS)
- L15 ANSWER 62 OF 62 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Effects of neuroleptics displaying antidepressant activity on behavior of rats in the forced swimming test
- AB Levomepromazine [60-99-1], thioridazine [50-52-2] and cis-chlorprothixene [113-59-7], neuroleptics with antidepressant activity, trans-chlorprothixene [4546-35-4], the therapeutically inactive isomer of chlorprothixene, clozapine [5786-21-0], an atypical neuroleptic, and imipramine [50-49-7], a classical antidepressant , were studied in the forced swimming test in rats after single or chronic administration. Levomepromazine (1.5 mg/kg), clozapine (2.5 and 5.0 mg/kg)mg/kg) and imipramine (10 mg/kg) after single administration, 1 h before the test, shortened the period of the immobility. After chronic administration only imipramine (10 mg/kg orally, twice daily, for 10 days) diminished the immobility. Levomepromazine, thioridazine, cis-chlorprothixene and trans-chlorprothixene (1.5 mg, orally, twice daily, for 10 days), 15-18 h after the last dose did not influence the immobility, although the behavioral parameters in the open field test were not depressed. Thus, the forced swimming test is not a suitable pharmacol. model for revealing antidepressant activities of certain neuroleptics that are useful in treating certain forms of human depression.
- AN 1985:534327 HCAPLUS <<LOGINID::20100601>>

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103:134327
DM
OREF 103:21309a,21312a
    Effects of neuroleptics displaying antidepressant activity on
     behavior of rats in the forced swimming test
    Gorka, Zbigniew; Janus, Krzysztof
ΑU
    Inst. Pharmacol., Pol. Acad. Sci., Krakow, 31-343, Pol.
CS
SO
    Pharmacology, Biochemistry and Behavior (1985), 23(2), 203-6
    CODEN: PBBHAU; ISSN: 0091-3057
DT
    Journal
LA English
OSC.G 7
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L2
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L3
             1 S SERTINDOLE/CN
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L4
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L5
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L6
L7
        104387 S DEPRESSION OR MDD OR DEPRESSIVE
L8
           843 S L6 AND L7
L9
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          1518 S L5 AND L9
L11
           457 S L7 AND L10
L12
            89 S L11 AND (PY<2003 OR AY<2003 OR PRY<2003)
L13
          9038 S SUICIDE OR SUICIDAL OR SUICIDALITY
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Connecting via Winsock to STN

Welcome to STN International! Enter x:X

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FILE 'HCAPLUS' ENTERED AT 11:44:41 ON 01 JUN 2010
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FULL ESTIMATED COST
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CA SUBSCRIBER PRICE
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         16193 D4
L16
         2105 DOPAMINE AND D4
=> s 19 and 116
         219 L9 AND L16
L17
=> s 117 and (PY<2002 or AY<2002 or PRY<2002)
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       4244485 AY<2002
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3713094 PRY<2002

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=> s dopamine (3a) d4 101729 DOPAMINE 16193 D4

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=> s 19 and 119 L20

179 L9 AND L19

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L21 75 L20 AND (PY<2002 OR AY<2002 OR PRY<2002)

=> s depression or antidepressant 99682 DEPRESSION

25756 ANTIDEPRESSANT

115310 DEPRESSION OR ANTIDEPRESSANT L22

=> s 121 and 122

3 L21 AND L22 L23

=> d 123 1-3 ti abs bib

- L23 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2010 ACS on STN
- Ziprasidone: a novel antipsychotic agent with a unique human receptor binding profile
- AΒ Ziprasidone is a novel antipsychotic agent with a unique combination of pharmacol. activities at human receptors. Ziprasidone has high affinity for human 5-HT receptors and for human dopamine D2 receptors. Ziprasidone is a 5-HT1A receptor agonist and an antagonist at 5-HT2A, 5-HT2C and

5-HT1B/1D receptors. Addnl., ziprasidone inhibits neuronal uptake of 5-HT and norepinephrine comparable to the antidepressant imipramine. This unique pharmacol. profile of ziprasidone may be related to its clin. effectiveness as a treatment for the pos., neg. and affective symptoms of schizophrenia with a low propensity for extrapyramidal side effects, cognitive deficits and weight gain. 2001:609740 HCAPLUS <<LOGINID::20100601>> 136:477 Ziprasidone: a novel antipsychotic agent with a unique human receptor binding profile Schmidt, A. W.; Lebel, L. A.; Howard, H. R.; Zorn, S. H. Groton Laboratories, CNS Discovery, Pfizer Global Research and Development, Groton, CT, 06340-1596, USA European Journal of Pharmacology (2001), 425(3), 197-201 CODEN: EJPHAZ; ISSN: 0014-2999 Elsevier Science B.V. Journal English OSC.G THERE ARE 83 CAPLUS RECORDS THAT CITE THIS RECORD (83 CITINGS) 83 RE.CNT 12 THERE ARE 12 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT L23 ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2010 ACS on STN Methods and compositions for the treatment of neuroleptic and related disorders using sertindole derivatives The invention relates to novel methods using, and pharmaceutical compns. and dosage forms comprising, sertindole derivs. Sertindole derivs. include, but are not limited to, nor-sertindole, 5-oxo-sertindole, dehydro-sertindole, and dehydro-nor-sertindole. The methods of the invention are directed to the treatment and prevention of neuroleptic and related disorders such as, but are not limited to, psychotic disorders, depression, anxiety, substance addiction, memory impairment and pain. For example, capsules were prepared containing a sertindole derivative 50.0 mg, lactose 48.5 mg, TiO2 0.5 mg, and Mg stearate 1.0 mg. 2000:861482 HCAPLUS <<LOGINID::20100601>> 134:32977 Methods and compositions for the treatment of neuroleptic and related disorders using sertindole derivatives Jerussi, Thomas P. Sepracor Inc., USA PCT Int. Appl., 33 pp. CODEN: PIXXD2 Patent English FAN.CNT 1 DATE APPLICATION NO. PATENT NO. KIND DATE _____ _____ ____ ______ _____ WO 2000072837 A2 20001207 WO 2000-US14984 20000531 <--АЗ WO 2000072837 20010517 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG US 6489341 B1 20021203 US 2000-580492 20000530 <--PRAI US 1999-137447P P 19990602 <--

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ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT

- OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)
- RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L23 ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Olanzapine: interaction study with imipramine
- AΒ Olanzapine is an "atypical" antipsychotic agent with a high affinity for serotonin 5HT2A/C, 5HT3, 5HT6, and dopamine D1, D2, D3, D4 receptors. Depressed patients with psychotic disorders frequently require treatment with concomitant antipsychotic and antidepressant medications. Imipramine pharmacokinetics serve as a marker for hepatic CYP2D6, CYP1A2, CYP3A activity. An open-label, three-way randomized crossover study was done to determine the safety, pharmacokinetics, and potential for a drug interaction between olanzapine (5 mg) and imipramine (75 mg). Each drug was administered alone and in combination. Nine healthy men, ages 32 to 54 yr, enrolled in the study. Psychomotor performance capacities, plasma olanzapine, imipramine, desipramine concns., and clin. laboratory tests were measured. Pharmacokinetic variables, vital signs, subjective tests for liveliness, and psychomotor outcomes were analyzed using a two-way ANOVA. Olanzapine was safe. Sedation, postural hypotension, and minor vital sign alterations occurred during all treatments. On the liveliness questionnaire, patients generally reported poorer (less lively) scores with olanzapine alone or coadministered with imipramine vs. baseline scores. These effects disappeared within 24 h after administration. Olanzapine alone and in combination decreased motor-speed tasks (finger tapping and visual-arm random reach) compared with base-line or imipramine treatment. Peak 6-h changes were statistically significant but clin. importance was only marginal. Olanzapine did not affect the kinetics of imipramine or desipramine and, therefore, did not show a metabolic drug interaction involving CYP2D6.
- AN 1997:748043 HCAPLUS <<LOGINID::20100601>>
- DN 128:57351
- OREF 128:11074h,11075a
- TI Olanzapine: interaction study with imipramine
- AU Callaghan, John T.; Cerimele, Benito J.; Kassahun, Kelem J.; Nyhart, Eldon H.; Hoyes-Beehler, Pamela J.; Kondraske, George V.
- CS Lilly Laboratory for Clinical Research, Department of Medicine and Pharmacology, Wishard Memorial Hospital and Indiana University Medical School, Indianapolis, IN, 46202, USA
- SO Journal of Clinical Pharmacology (1997), 37(10), 971-978 CODEN: JCPCBR; ISSN: 0091-2700
- PB Lippincott-Raven
- DT Journal
- LA English
- OSC.G 15 THERE ARE 15 CAPLUS RECORDS THAT CITE THIS RECORD (15 CITINGS)
- RE.CNT 18 THERE ARE 18 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT